

Atty. Docket No.:

8039/1090

TECH CENTER 1600/2000

Application of:

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE Riechmann, et al.

Serial No.:

09/710,444

Filed:

November 10, 2000

Entitled:

Selection System

Examiner:

B. Celsa

Group Art Unit:

1639

Conf. No.:

5253

CERTIFICATE OF MAILING UNDER 37 C.F.R. § 1.8a

I hereby certify that this correspondence (and any paper or fee referred to as being enclosed) is being deposited with the United States Post Office as First Class Mail on the date indicated below in an envelope addressed to Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

Kathleen Williams

Name of Person Mailing Paper

Signature of Person Mailing Paper

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

REQUEST UNDER 37 C.F.R. §1.181 TO WITHDRAW HOLDING OF ABANDONMENT

Sir:

Applicants hereby request withdrawal of the holding of abandonment announced in the Notice of Abandonment mailed September 22, 2003.

This is filed in response to the Notice of Abandonment of Non-Provisional Application mailed by the US Patent & Trademark Office on September 22, 2003.

The Notice stated that Applicants failed to comply with the Sequence Rules within 6 months of the first letter, dated February 27, 2002, and within 1 month of the Non-bona fide response letter dated March 17, 2003. Applicants submit that bona fide responses to both communications from the Patent Office were timely filed and that the holding of abandonment is in error. This assertion is supported as follows. Enclosed are the following:

- 1) A copy of the Notice to Comply mailed February 27, 2002;
- 2) A copy of the response filed July 25, 2002, including fee and petition for Extension of Time. This response included paper and diskette copies of the sequence listing and the required statement under 37 CFR §1.821 (f) and (g);
- 3) A copy of the second Notice to Comply (Non-bona fide). That notice stated that the disk was melted by the U.S.P.T.O. and set a 1 month non-extendable deadline for response. Applicants note that the response was properly mailed to the Arlington, VA address established for Sequence Diskette Submission to avoid irradiation damage to the diskette;
- 4) A copy of the filing of April 8, 2003, responsive to the March 17,2003 communication. That filing included a new diskette, paper copy of the Sequence Listing and Statement under 37 C.F.R. 1.821 (f) and (g) and a return postcard. Applicants submit that the April 8, 2003 filing was well within the 1 month time set in the second Notice to Comply; and

A copy of the O I.P.E.-stamped copy of the return postcard filed with the April 8, 2003 response to the second Notice to Comply. The filing was stamped by the U.S.P.T.O. on April 11, 2003. The return of the date-stamped postcard acknowledges receipt of all items asked for in the second Notice to Comply: a substitute diskette; a substitute paper copy of the Sequence Listing; and a Statement Under 37 CFR 1.821 (f) and (g).

In view of the above, Applicants submit that Applicants have fully and timely responded to all Notices from the U.S.P.T.O. Applicants therefore respectfully request notice that the holding of abandonment has been withdrawn.

Because it is believed that the Notice of Abandonment is in error, it is believed that no fees are due with this filing. However, the Commissioner for Patents is authorized to charge all fees in the total amount to Deposit Account No. 16-0085, Reference 8039/1090.

Date: 9\16\103

Respectfully submitted,

Name: Kathleen Williams Registration No.: 34,380 Customer No.: 29933 Palmer & Dodge LLP

111 Huntington Avenue Boston, MA 02199-7613

Tel: 617-239-0100



Atty. Docket No.:

8039/1090

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of:

Riechmann, et al.

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PECHOENT 3 2003
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Kathleen Williams

Name of Person Mailing Paper

Signature of Person Mailing Paper

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

TRANSMITTAL LETTER

Enclosed for filing the above-identified patent application, please find the following documents:

- 1. Request Under 37. C.F.R. §1.181 to Withdraw Holding of Abandonment;
- Copy of the Office Action Mailed February 27, 2002; 2.
 - Copy of the Response to the 2/27/02 Office Action (filed 7/25/02).
- 4. Copy of the Office Action mailed 3/17/03;
- Copy of the Response to the 3/17/03 Office Action (filed April 8, 2003). 5.
- 6. Copy of the O.I.P.E.-stamped copy of the Return Post Card filed with the April 8, 2003 Response to the Second Notice to Comply; and
- Return Post Card. 7.

The Commissioner for Patents is hereby authorized to charge any additional fees or credit any overpayment in the total fees to Deposit Account No. 16-0085, Reference 8039/1090. A duplicate of this transmittal letter is enclosed for this purpose.

Date:

9126103

Respectfully submitted,

Name: Kathleen Williams Registration No.: 34,380 Customer No.: 29933

Palmer & Dodge LLP 111 Huntington Avenue Boston, MA 02199-7613

Tel: 617-239-0100

Serial No. 09 710, 444 File No. 8 Applicant(s): Rigenman, et al.	034 1090 By: Kmy
Title: Selection System	
The Following, DUE in the USPTO, was received [] Cert. of Mailing by Express Mail (37 CFR 1.10) Express Mail Label No	by the PTO Mail Room on the date stamped hereon: Response to Notice to File Missing Parts Oppy of Part 2 of NFMP
Cert. of Mailing under 37 CFR 1.8(a) Patent Application (total pgs) Provisional or [] Non-Provision (pgs) Specification (pgs) Abstract, (pgs) Claims (# claims New Patent Application Transmitta Provisional Patent Application Cover Sheet 1 1 2003	Diskette Containing Nucleotide and/or Amino Acid Sequence Listing Substitute Priority Document(s) # Amendment/Response Petition for Extension of Time (x2) Check in the amount of Check #
[] Application Data Sheet [] Drawings Sheet(s) (FIGS [] Formal or [] Assignment of [] Recordation Cover Sheet Form PTO-1595 [] Information Disclosure Statement [] Form PTO 1449 and Copies of Cited References	Transmittal of Formal Drawings Motion/Opposition/Reply Request for Cont'd Examination (RCE) Notice of Appeal Appeal Brief (x3) Issue Fee Transmittal Transmittal Letter (x2)
of Notice to Compy, Statement Under MAILED April 8	37 CFR 1.831 (f) === (6) 4

RECEIVED

VAPR 1 7 2003

PATENT DEPT.



United States Patent and Trademark Office

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER OF PATENTS AND TRADEMARKS Washington, D.C. 20231 www.uspto.gov

FIRST NAMED INVENTOR APPLICATION NO. FILING DATE ATTORNEY DOCKET NO. CONFIRMATION NO. 09/710,444 11/10/2000 Lutz Riechmann 8654/1090 5253 29933 7590 03/17/2003 PALMER & DODGE, LLP EXAMINER KATHLEEN M. WILLIAMS CELSA, BENNETT M 111 HUNTINGTON AVENUE BOSTON, MA 02199 ART UNIT PAPER NUMBER 1639 DATE MAILED: 03/17/2003

Please find below and/or attached an Office communication concerning this application or proceeding.

Response Due Notice to Complete Response Due Notice Due Notice Response Due Notice Due Notic



UNITED STATES DEPARTMENT OF COMMERCE Patent and Trademark Office COMMISSIONER OF PATENTS AND TRADEMARKS Washington, D.C. 20231

SERIAL NUMBER	FILING DATE	FIRST NAMED APPLICANT	ATTORNEY DOCKET NO.
09/ 710,444			

EXAMINER

ART UNIT PAPER NUMBER

1639 13

Please find below a communication from the EXAMINER in charge of this application

NOTICE TO COMPLY: SEQUENCE RULES (NONBONAFIDE)

1. Applicant's submission of a computer readable form (CRF) and corresponding paper sequence listing in paper no. 12(dated 12/10/02) is acknowledged.

However, this application fails to comply with the requirements of 37 CFR 1.821 through 1.825 for the reason(s) set forth on the attached Notice To Comply With Requirements For Patent Applications Containing Nucleotide Sequence And/Or Amino Acid Sequence Disclosures and CRF PROBLEM REPORT. Applicant must comply with the requirements of the sequence rules (37 CFR 1.821 - 1.825) before the application can be examined under 35 U.S.C. \$\$ 131 and 132.

Applicant is given ONE MONTH (NON-EXTENDIBLE) from the mailing date of this communication within which to <u>COMPLY WITH</u>

THE SEQUENCE RULES, 37 CFR 1.821 - 1.825. Failure to comply with these requirements will result in ABANDONMENT of the application. Direct the reply to the undersigned. Applicant is requested to return a copy of the attached Notice to Comply with the reply.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Examiner Celsa whose telephone number is (703) 305-7556). If the examiner cannot be reached, inquiries can be directed to Supervisory Patent Examiner Andrew Wang whose telephone number is (703) 306-3217. Inquiry of a general nature or relating to the status of this application should be directed to the Group receptionist whose telephone number is (703) 308-0196.

Primary Examiner Celsa ART UNIT 1639

March 14, 2003

BENNETT CELSA
PRIMATY EXAMPLE

////

Application No.: L	⁾ 710,444

NOTICE TO COMPLY WITH REQUIREMENTS FOR PATENT APPLICATIONS CONTAINING NUCLEOTIDE SEQUENCE AND/OR AMINO ACID SEQUENCE DISCLOSURES

Applicant must file the items indicated below within the time period set the Office action to which the Notice is attached to avoid abandonment under 35 U.S.C. § 133 (extensions of time may be obtained under the provisions of 37 CFR 1.136(a)).

The nucleotide and/or amino acid sequence disclosure contained in this application does not comply with the requirements for such a disclosure as set forth in 37 C.F.R. 1.821 - 1.825 for the following reason(s):

X	1. This application clearly fails to comply with the requirements of 37 C.F.R. 1.821-1.825. Applicant's attention is directed to the final rulemaking notice published at 55 FR 18230 (May 1, 1990), and 1114 OG 29 (May 15, 1990). If the effective filing date is on or after July 1, 1998, see the final rulemaking notice published at 63 FR 29620 (June 1, 1998) and 1211 OG 82 (June 23, 1998).
	. This application does not contain, as a separate part of the disclosure on paper copy, a "Sequence Listing" as required by 37 C.F.R. 1.821(c).
	3. A copy of the "Sequence Listing" in computer readable form has not been submitted as required by 37 C.F.R. 1.821(e).
	A copy of the "Sequence Listing" in computer readable form has been submitted. However, the content of the computer readable form does not comply with the requirements of 37 C.F.R. 1.822 and/or 1.823, as indicated on the attached copy of the marked -up "Raw Sequence Listing."
X	5. The computer readable form that has been filed with this application has been found to be damaged and/or unreadable as indicated on the attached CRF Diskette Problem Report. A Substitute computer readable form must be submitted as required by 37 C.F.R. 1.825(d).
	6. The paper copy of the "Sequence Listing" is not the same as the computer readable from of the "Sequence Listing" as required by 37 C.F.R. 1.821(e).
	. Other:
App	licant Must Provide:
X	An initial or substitute computer readable form (CRF) copy of the "Sequence Listing".
x	An initial or substitute paper copy of the "Sequence Listing", as well as an amendment directing its entry into the specification.
X	A statement that the content of the paper and computer readable copies are the same and, where applicable, include no new matter, as required by 37 C.F.R. 1.821(e) or 1.821(f) or 1.821(g) or 1.825(b) or 1.825(d).
For	questions regarding compliance to these requirements, please contact:
For	Rules Interpretation, call (703) 308-4216 CRF Submission Help, call (703) 308-4212 entIn Software Program Support
	Technical Assistance
•	PLEASE RETURN A COPY OF THIS NOTICE WITH YOUR REPLY

CRF Problem Report

BIOJECHNOLOGY DEMS

The Scientific and Technical Information Center (STIC) experienced a problem when processing the following computer readable form (CRF):

Application Serial Number: 09 710, 444A
Filing Date: 1120 2000
Date Processed by STIC: 1129/02

STIC Contact: Mark Spencer, 703-308-4212

RECEIVED

DEC 1 0 2002

Nature of Problem:

TECH CENTER 1600/2900

n 4	[COITOL
The CRF (was):	
- Melted	•
(circle one) Damaged or Unreadable (for	or Unreadable, see attached)
Blank (no files on CRF) (see attached)	,
Empty file (filename present, but no by	tes in file) (see attached)
Virus-infected. Virus name:	The STIC will not process the CRF.
Not saved in ASCII text	•
Sequence Listing was embedded in the	file. According to Sequence Rules,
submitted file should only be the Seq	
Did not contain a Sequence Listing. (s	ee attached sample)
Other:	

PLEASE USE THE CHECKER VERSION 3.1 PROGRAM TO REDUCE ERRORS. SEE BELOW FOR ADDRESS:

http:/www.uspto.gov/web/offices/pac/checker

Applicants submitting genetic sequence information electronically on diskette or CD-Rom should be aware that there is a possibility that the disk/CD-Rom may have been affected by treatment given to all incoming mail. Please consider using alternate methods of submission for the disk/CD-Rom or replacement disk/CD-Rom. Any reply including a sequence listing in electronic form should NOT be sent to the 20231 zip code address for the United States Patent and Trademark Office, and instead should be sent via the following to the indicated addresses:

- 1. EFS-Bio (<http://www.uspto.gov/ebc/efs/downloads/documents.htm>, EFS Submission User Manual ePAVE)
- 2. U.S. Postal Service: U.S. Patent and Trademark Office, Box Sequence, P.O. Box 2327, Arlington, VA 22202

3. Hand Carry directly to:

- U.S. Patent and Trademark Office, Technology Center 1600, Reception Area, 7th Floor, Examiner Name, Sequence Information, Crystal Mall One, 1911 South Clark Street, Arlington, VA 22202
- U.S. Patent and Trademark Office, Box Sequence, Customer Window, Lobby, Room 1B03, Crystal Plaza Two, 2011 South Clark Place, Arlington, VA 22202
- Federal Express, United Parcel Service, or other delivery service to: U.S. Patent and Trademark Office, Box Sequence, Room 1B03-Mailroom, Crystal Plaza Two, 2011 South Clark Place, Arlington, VA 22202

Revised 01/29/2002



UNITED STATES PATENT AND TRADEMARK OFFICE

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER OF PATENTS AND TRADEMARKS Washington, D.C. 20231 www.uspto.gov

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/710,444	11/10/2000	Lutz Riechmann	8654/1090	5253
7:	590 02/27/2002	OIPE		
Palmer & Doo		4	EXAMI	NER ·
One Beacon St Boston, MA 0		SEP 2 9 2003	CELSA, BEI	NNETT M
		M A	ART UNIT	PAPER NUMBER
		E DOEMAGE CE	1627	
		EMB	DATE MAILED: 02/27/2002	

Please find below and/or attached an Office communication concerning this application or proceeding.

neary

Palmer & Dodge LLP Drop
Patent Department

MAR 1 4 2802
Palmer & Dedge, LLP
Patent Department



UNITED STALES DEPARTMENT OF COMMERCE Patent and Trademark Office
COMMISSIONER OF PATENTS AND TRADEMARKS Washington, D.C. 20231

SERIAL NUMBER	FILING DATE	FIRST NAMED APPLICANT	ATTORNEY DOCKET NO.
09/ 710,444	·		

EXA	MINER
ART UNIT	PAPER NUMBER
1627	5

Please find below a communication from the EXAMINER in charge of this application

Sequence Rule Compliance: NOTICE TO COMPLY

This application fails to comply with the sequence rule quirements of 37 C.F.R. §§ 1.821-1.825 for the reason(s) set forth on the attached Notice To Comply With Requirements For Patent Applications Containing Nucleotide Sequence And/Or Amino Acid Sequence Disclosures. This application encompasses sequences needing sequence identifiers (e.g. see pages 9, 15, 24, 30, 35, 36, 38, figures etc.).

APPLICANT IS GIVEN 30 days FROM THE DATE OF THIS LETTER WITHIN WHICH TO COMPLY WITH THE SEQUENCE RULES, 37 C.R.F. §§ 1.821-1.825. Failure to comply with these requirements will result in ABANDONMENT of the application under 37 C.F.R. § 1.821(g). Extensions of time may be obtained by filing a petition accompanied by the extension fee under the provisions of 37 C.F.R. § 1.136. In no case may an applicant extend the period for response beyond the six month statutory period. Direct the response to the undersigned. Applicant is requested to return a copy of the attached Notice to Comply with the response.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Examiner Celsa whose telephone number is (703) 305-7556 If the examiner cannot be reached, inquiries can be directed to Supervisory Patent Examiner Venkat whose telephone number is (703) 308-0570. The fax number for the organization where this application or proceeding is assigned is (703) 308-4242. Any inquiry of a general nature or relating to the status of this application should be directed to the Group receptionist whose telephone number is (703) 308-0196.

Bennett Celsa (au 1627) (Feb. 25,2002)

BENNETT CELSA PRIMADY EXAMINED

M

Application No. 09/710, 444

NOTICE TO COMPLY WITH REQUIREMENTS FOR PATENT APPLICATIONS CONTAINING NUCLEOTIDE SEQUENCE AND/OR AMINO ACID SEQUENCE DISCLOSURES

for such a disclosure as set forth in 37 CFR 1.821 - 1.825 for the following reason(s):	with the require
1. This application clearly fails to comply with the requirements of 37 CFR 1.821 - 1.825. App directed to these regulations, published at 1114 OG 29, May 15, 1990 and at 55 FR 18230, Max	plicant's attention is ay 1, 1990.
2. This application does not contain, as a separate part of the disclosure on paper copy, a "Secretarized by 37 CFR 1.821(c).	quence Listing ^{*,} as
3. A copy of the "Sequence Listing" in computer readable form has not been submitted as requi	
as required as required the submitted as required as r	ired by 37 CFR 1.821
4. A copy of the "Sequence Listing" in computer readable form has been submitted. However, to computer readable form does not comply with the requirements of 37 CFR 1.822 and/or 1.823, a attached marked-up copy of the "Raw Sequence Listing."	
5. The computer readable form that has been filed with this application has been found to be dar unreadable as indicated on the attached CRF Diskette Problem Report. A substitute computer resubmitted as required by 37 CFR 1.825(d).	maged and/or eadable form must be
6. The paper copy of the "Sequence Listing" is not the same as the computer readable form of the as required by 37 CFR 1.821(e).	
7. Other:	
Applicant must provide:	• •

An initial or substitute computer readable form (CRF) copy of the "Sequence Listing"

An initial or substitute paper copy of the "Sequence Listing", as well as an amendment directing its entry into the

A statement that the content of the paper and computer readable copies are the same and, where applicable, include no new matter, as required by 37 CFR 1.821(e) or 1.821(f) or 1.821(g) or 1.825(b) or 1.825(d)

For questions regarding compliance with these requirements, please contact:

For Rules Interpretation, call (703) 308-1123 For CRF submission help, call (703) 308-4212 For Patentin software help, call (703) 308-6856

Please return a copy of this notice with your response.

Attachment for PTO-948 (Rev. 03/01, or earlier) 6/18/01

The below text replaces the pre-printed text under the heading, "Information on How to Effect Drawing Changes," on the back of the PTO-948 (Rev. 03/01, or earlier) form.

INFORMATION ON HOW TO EFFECT DRAWING CHANGES

1. Correction of Informalities - 37 CFR 1.85

New corrected drawings must be filed with the changes incorporated therein. Identifying indicia, if provided, should include the title of the invention, inventor's name, and application number, or docket number (if any) if an application number has not been assigned to the application. If this information is provided, it must be placed on the front of each sheet and centered within the top margin. If corrected drawings are required in a Notice of Allowability (PTOL-37), the new drawings MUST be filed within the THREE MONTH shortened statutory period set for reply in the Notice of Allowability. Extensions of time may NOT be obtained under the provisions of 37 CFR 1.136(a) or (b) for filing the corrected drawings after the mailing of a Notice of Allowability. The drawings should be filed as a separate paper with a transmittal letter addressed to the Official Draftsperson.

2. Corrections other than Informalities Noted by Draftsperson on form PTO-948.

All changes to the drawings, other than informalities noted by the Draftsperson, MUST be made in the same manner as above except that, normally, a highlighted (preferably red ink) sketch of the changes to be incorporated into the new drawings MUST be approved by the examiner before the application will be allowed. No changes will be permitted to be made, other than correction of informalities, unless the examiner has approved the proposed changes.

Timing of Corrections

Applicant is required to submit the drawing corrections within the time period set in the attached Office communication. See 37 CFR 1.85(a).

Failure to take corrective action within the set period will result in **ABANDONMENT** of the application.



Atty. Docket No.:

8039/1090

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of:

Riechmann, et al.

Serial No.:

09/710,444

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Mary Wilson

Name of Person Mailing Paper

Signature of Person Mailing Paper

U.S. Patent & Trademark Office

Box: Sequence PO Box 2327

Arlington, VA 22202

TRANSMITTAL LETTER

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- 1. Response to Office Action mailed March 17, 2003;
- 2. Copy of Notice to Comply;
- 3. Substitute Paper Copy of the Sequence Listing;
- 4. Substitute Computer Readable Copy of the Sequence Listing;
- 5. Statement Under 37 C.F.R. §1.821(f) and (g); and
- 6. Return Post Card.

It is believed that no fees are due. However, if necessary, the Commissioner for Patents is hereby authorized to charge all fees in the total amount to Deposit Account No. 16-0085, Reference 8039/1090. A duplicate of this transmittal letter is enclosed for this purpose.

Respectfully submitted,

Date

April 8, 2003

ال يسيعه

Response due __

Statutory period

Palmer & Dodge LLP
Patent Department

1 1 (

Name: Mark J. FitzGerald Registration No.: 45,928 Palmer & Dodge LLP

111 Huntington Avenue Boston, MA 02199-7613

Tel: 617-239-0100



Atty. Docket No.:

8039/1090

PATENT

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Mary Wilson

Name of Person Mailing Paper

Signature of Person Mailing Paper

U.S. Patent & Trademark Office

Box: Sequence PO Box 2327

Arlington, VA 22202

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Respectfully submitted,

Date: April 8, 2003

Name: Mark J. FitzGerald

Registration No.: 45,928 Palmer & Dodge LLP

111 Huntington Avenue Boston, MA 02199-7613

Tel: 617-239-0100

Application	No.: L	0,444
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NOTICE TO COMPLY WITH REQUIREMENTS FOR PATENT APPLICATIONS CONTAINED NUCLEOTIDE SEQUENCE AND/OR AMINO ACID SEQUENCE DISCLOSURES

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The nucleotide and/or amino acid sequence disclosure contained in this application does not comply with the requirements for such a disclosure as set forth in 37 C.F.R. 1.821 - 1.825 for the following reason(s):

X	1. This application clearly fails to comply with the requirements of 37 C.F.R. 1.821-1.825. Applicant's attention is directed to the final rulemaking notice published at 55 FR 18230 (May 1, 1990), and 1114 OG 29 (May 15, 1990). If the effective filing date is on or after July 1, 1998, see the final rulemaking notice published at 63 FR 29620 (June 1, 1998) and 1211 OG 82 (June 23, 1998).
	. This application does not contain, as a separate part of the disclosure on paper copy, a "Sequence Listing" as required by 37 C.F.R. 1.821(c).
	3. A copy of the "Sequence Listing" in computer readable form has not been submitted as required by 37 C.F.R. 1.821(e).
	. A copy of the "Sequence Listing" in computer readable form has been submitted. However, the content of the computer readable form does not comply with the requirements of 37 C.F.R. 1.822 and/or 1.823, as indicated on the attached copy of the marked -up "Raw Sequence Listing."
X	5. The computer readable form that has been filed with this application has been found to be damaged and/or unreadable as indicated on the attached CRF Diskette Problem Report. A Substitute computer readable form must be submitted as required by 37 C.F.R. 1.825(d).
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App	olicant Must Provide:
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X	A statement that the content of the paper and computer readable copies are the same and, where applicable, include no new matter, as required by 37 C.F.R. 1.821(e) or 1.821(f) or 1.821(g) or 1.825(b) or 1.825(d).
For	questions regarding compliance to these requirements, please contact:
For	Rules Interpretation, call (703) 308-4216
For	CRF Submission Help, call (703) 308-4212
Pat	tentin Software Program Support
	Technical Assistance
	To Purchase Patentin Software703-306-2600

PLEASE RETURN A COPY OF THIS NOTICE WITH YOUR REPLY



Atty. Docket No.:

8039/1090

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of:

Riechmann, et al.

Serial No.:

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Mary Wilson

Name of Person Mailing Paper

Signature of Person Mailing Paper

U.S. Patent and Trademark Office

Box: Sequence

P.O. Box 2327

Arlington, VA 22202

RESPONSE TO OFFICE ACTION

Sir:

This is filed in response to the Office Action of Non-Provisional Application mailed March 17, 2003 in the above-noted U.S. patent application.

The Office Action stated that the Applicant must provide substitute computer readable and paper copies of the sequence listing along with the sequence listing statement required under 37 C.F.R. §1.821(f) and (g), because the disk submitted December 10, 2002 was damaged by the P.T.O.

Enclosed please find a Substitute paper copy of the Sequence Listing, a Substitute Computer Readable copy of the Sequence Listing, and the Statement Under 37 C.F.R. §1.821(f) and (g).

It is believed that no fees are due. However, if necessary, the Commissioner for Patents is hereby authorized to charge all fees in the total amount to Deposit Account No. 16-0085, Reference 8039/1090. A duplicate of this transmittal letter is enclosed for this purpose.

Respectfully submitted,

Date: April 8, 2003

Name: Mark J. FitzGerald Registration No.: 45,928 Palmer & Dodge LLP 111 Huntington Avenue Boston, MA 02199-7613

Tel: 617-239-0100

SEP 2 9 2003 BY

Atty. Docket No.:

8039/1090

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of:

Riechmann, et al.

Serial No.:

09/710,444

Filed:

November 10, 2000

Entitled:

"Selection System"

Examiner:

B. Celsa

Group Art Unit:

1639

Conf. No.:

5253

CERTIFICATE OF MAILING UNDER 37 C.F.R. § 1.8a

I hereby certify that this correspondence (and any paper or fee referred to as being enclosed) is being deposited with the United States Post Office as First Class Mail on the date indicated below in an envelope addressed to: U.S. Patent & Trademark Office, Box: Sequence, P.O. Box 2327, Arlington, VA 22202.

Mary Wilson

Name of Person Mailing Paper

Signature of Person Mailing Paper

U.S. Patent & Trademark Office

Box: Sequence PO Box 2327

Arlington, VA 22202

STATEMENT UNDER 37 C.F.R. §1.821(f) and (g)

Sir:

This paper is submitted in response to the Office Action mailed by the USPTO on March 17, 2003.

In accordance with 37 C.F.R. §1.821 (f) I hereby state that the paper copy and the computer readable form of the Sequence Listing submitted herewith in the above-identified patent application are supported in the application and contain no new matter. I hereby state that the information recorded in computer readable form is identical to the written sequence listing.

In accordance with 37 C.F.R. §1.821 (g), I hereby state that the computer readable form of the Sequence Listing submitted herewith contains no new matter.

Respectfully submitted,

Date: April 8, 2003

Name: Mark J. FitzGerald Registration No.: 45,928 Palmer & Dodge LLP 111 Huntington Avenue

Boston, MA 02199-7613

Tel: 617-239-0100



SEQUENCE LISTING

- <110> Riechmann, Lutz

 Kristensen, Peter

 Jestin, Jean-Luc

 Winter, Gregory
- <120> Selection System
- <130> 8039/1090
- <140> 09/710,444
- <141> 2000-11-20
- <150> GB 9810223.9
- <151> 1998-05-13
- <150> GB 9810228.8
- <151> 1998-05-13
- <150> PCT/GB99/01526
- <151> 1999-05-13
- <160> 79
- <170> PatentIn version 3.1

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A, T or C

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<221> misc_feature

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                                                                     60
ttgsyaayrs yasyasyagb nttgttatta ctcsyanycv nncygdccat ggcccaggtg
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                                                                      50
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aagettgeat geaceagete tdteaaggag acagteataa tgaggegget
                                                                 50
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<222> (38)..(38)
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<221> misc_feature

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                                                                     55
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                                      10
                                                          15
   Ala Gln Pro Ala Met Ala
               20
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1
                                  10
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Pro Thr Gln Pro Ala Met Ala

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                                                                     50
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                                                                     50
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                                                                      50
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        50
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<212> PRT

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                5
                                   10
                                                      15
Ala Gln Pro Ala Met Ala
            20
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                                                15
Val Gln Pro Ala Met Ala
          20
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<221> misc_feature
<222> (1)..(30)
<223> Synthetic oligonucleotide primer used as substrate for Thermus aq
      uaticus DNA polymerase I
<400> 49
aaatacaaca ataaaacgcc acatcttgcg
                                                                      30
<210> 50
<211> 20
<212> DNA
<213> Artificial sequence
<220>
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<223> Synthetic oligonucleotide sequence insert containing PstI restric

)

tion site and frame shift for H102A mutant barnase fusion construct fused to p3 gene of phage fd-3.

<220>

<221> misc_feature

<222> (1)..(20)

<223> Synthetic oligonucleotide sequence insert containing PstI restric tion site and frame shift for H102A mutant barnase fusion construct fused to p3 gene of phage fd-3.

<400> 50

ctgcaggcgg tgcggccgca

20

<210> 51

<211> 24

<212> DNA

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<220>

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<220>

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<223> Synthetic oligonucleotide used for random priming

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<223> n at position 19 can be G, A, T or C.
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<221> misc_feature
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<220>
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<221> misc_feature
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<210> 52
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      i genomic DNA sequences.
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<400> 52
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cgtgcgagcc tgcagagctc agg

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<400> 53

Leu Gln Ser Ser Gly Asp Cys Val Ile Ser Asp Thr Cys Ile Ala Gly 1 5

10

15

Met Ala Glu Ala Ala Cys Glu Glu Lys Phe Ser Ser Gln Asn Val

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Gly Leu Thr Ile Thr Val Thr Pro Cys Leu Ser Ser Ala
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40

45

<210> 54

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<222> (1)..(44)

<223> Barstar binding barnase-p3 fusion insert

<400> 54

Leu Gin Ser Ser Gly Cys Gly Ser Ser Gly Ser Ser Ile Asn Cys Leu

1 10 15

Pro Cys Gly Ala Thr Ser Arg Gly Thr Ser Pro Leu Ala Ser Gly Leu

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Pro Ser Ser Ala Thr Ile His Cys Leu Ser Ser Ala
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40

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<223> Barstar binding barnase-p3 fusion insert

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Leu Gln Ser Ser Gly Asp Ser Ala Gly Cys Lys Asn Met Thr Gly Gly

1

5

10

15

Arg Leu Tyr Ala His Thr Leu Glu Ala Ile Ile Pro Gly Phe Ala Val

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Ser Ala Pro Ala Cys Glu Pro Ala
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40

<210> 56

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<212> PRT

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<400> 56

Leu Gln Ser Ser Gly Cys Val Arg Leu Lys Arg Thr Ser Val Asn His

1 10 15

Gln Pro Asp Ala Trp Pro Glu Pro His Leu Lys Ala Ala Cys Glu Pro

Ala

<210> 57

<211> 44

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<400> 57

Leu Gln Ser Ser Gly Cys Gly Ser Ser Gly Ser Ser Ile Asn Cys Leu

1 5 10 15

Pro Cys Gly Ala Thr Ser Arg Gly Thr Ser Pro Leu Ala Ser Gly Leu

20

25

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Pro Ser Ser Ala Thr Val Gln Cys Leu Ser Ser Ala
```

40

<210> 58

<211> 41

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<222> (1)..(41)

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Leu Gln Ser Ser Gly Lys Ile Val Gln Ala Gly Ala Asn Ile Gln Asp

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Gly Cys Ile Met His Gly Tyr Cys Asp Thr Asp Thr Ile Val Gly Glu

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Asn Gly His Ile Gly Leu Ser Ser Ala
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<211> 45

<212> PRT

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Leu Gln Ser Ser Gly Val Cys Val Ile Ser Asp Thr Cys Ile Ala Gly

1 10 15

Thr Ala Glu Ala Ala Cys Glu Glu Lys Phe Ser Ser Gln Asn Val

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Gly His Thr Ile Thr Glu Thr Pro Cys Leu Ser Ser Ala
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45

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Pro Cys Gly Ala Thr Ser Arg Gly Thr Ser Pro Leu Ala Ser Gly Leu

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Pro Ser Ser Ala Thr Ile Gln Cys Leu Ser Ser Ala
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<210> 61

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<223> Barstar binding barnase-p3 fusion insert

<400> 61

Leu Gln Ser Ser Gly Gln Asp Ser Gln Arg Glu His Ala Ser His Thr

1 5 10 15

Ala Glu Asp Asp Cys Glu Asp Gln Thr Arg Ile His Gln His Ile Arg

```
Glu Val Asp Phe Val Asp Thr Pro Gln Glu Val Asp Asp Cys Arg Ala

35 40 45
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Ala Leu Ser Ser Ala

50

<210> 62

<211> 33

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<400> 62

Leu Gln Ser Ser Gly Cys Val Arg Leu Lys Arg Thr Ser Val Asn His

1

5

10

```
Gln Pro Asp Ala Trp Pro Glu Pro His Leu Lys Ala Ala Cys Glu Pro
            20
                                25
```

Ala

<210> 63

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<222> (1)..(9)

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<400> 63

Leu Gln Ser Ser Gly Val Arg Pro Ala

<210> 64

<211> 44

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<222> (1)..(44)

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<400> 64

Leu Gln Ser Ser Gly Cys Gly Ser Ser Gly Ser Ser Ile Asn Cys Leu

1 5 10 15

Pro Cys Gly Ala Thr Ser Arg Gly Thr Ser Pro Leu Ala Ser Gly Leu
20 25 30

Pro Ser Ser Ala Thr Ile Gln Cys Leu Ser Ser Ala

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1
                5
                                    10
                                                        15
Thr Asn Asp Arg Asp Phe Thr His Thr Pro Leu Ser Ser Ala
            20
                                25
                                                    30
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<210> 66

<210> 65

<211> 36

<212> PRT

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               5
                                 10
                                                        15
Thr Ala Thr Asn Ala Val Leu Ser Ala Asp Ser Leu Ser Leu Gly Gly
           20
                                25
                                                    30
Gly Glu Pro Ala
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<210> 67

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1
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              5
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Ser Ser Ala
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                5
                                    10
                                                       15
Thr Ala
<210> 69
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<222> (1)..(9)

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<400> 69

Leu Gln Ser Ser Gly Val Arg Pro Ala

1 5

<210> 70

<211> 36

<212> PRT

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<221> MISC_FEATURE

<222> (1)..(36)

<223> Barstar binding barnase-p3 fusion insert

<400> 70

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1

5

10

```
Glu Ala Pro Val Ala Lys Ala Glu Ala Lys Pro Glu Thr Pro Ala His
20 25 30
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Leu Ser Ser Ala

35

<210> 71

<211> 33

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<223> Barstar binding barnase-p3 fusion insert.

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<221> MISC_FEATURE

<222> (1)..(33)

<223> Barstar binding barnase-p3 fusion insert

<400> 71

Leu Gln Ser Ser Gly Cys Val Arg Leu Lys Arg Thr Ser Val Asn His

1

5

10

```
Gln Pro Asp Ala Trp Pro Glu Pro His Leu Lys Ala Ala Cys Glu Pro
20 25 30
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Ala

<210> 72

<211> 36

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<220>

<221> MISC_FEATURE

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<223> Barstar binding barnase-p3 fusion insert

<400> 72

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1

5

10

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Asp Ser Ile Gly Ala Tyr Leu Phe Val Asp Met Ala His Val Ala Ala
20 25 30
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Leu Ser Ser Ala

35

<210> 73

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<223> Vector pK1 polylinker sequence.

<220>

<221> misc_feature

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<223> Vector pK1 polylinker sequence

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aatgctggcg gcggcccagc cggcctttct gaggggtcga ctatagaagg acgaggggcc 60

cacgaaggag gtggggtacc cggttccgag ggtggttccg gttccggtga ttttgat

<210> 74

<211> 39

<212> PRT

<213> Artificial sequence

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<220>

<221> MISC_FEATURE

<222> (1)..(39)

<223> Polypeptide encoded by pK1 vector polylinker sequence

<400> 74

Asn Ala Gly Gly Pro Ala Gly Leu Ser Glu Gly Ser Thr Ile Glu

1 5 10 15

Gly Arg Gly Ala His Glu Gly Gly Gly Val Pro Gly Ser Glu Gly Gly
20 25 30

Ser Gly Ser Gly Asp Phe Asp

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<210> 75
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                                                                   60
acgaagcagc tggggtaccg gttccgaggg tggttccggt tccggtgatt ttgatta
                                                                  117
<210> 76
<211> 39
<212> PRT
<213> Artificial sequence
                         - . .
<220>
```

<223> Polypeptide sequence encoded by vector pK2 polylinker region.

)

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<222>
      (1)..(39)
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<220>
<221> MISC_FEATURE
<222> (38)..(38)
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<220>
<221> MISC_FEATURE
<222> (36)..(36)
<223> X represents a stop codon (TGA)
<400> 76
Asn Ala Gly Gly Pro Ala Gly Leu Ser Glu Gly Ser Thr Ile Glu
1
               5
                                   10
                                                      15
```

Gly Arg Gly Pro Thr Lys Gln Leu Gly Tyr Arg Phe Arg Gly Trp Phe . . .

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Arg Phe Arg Xaa Phe Xaa Leu
        35
<210> 77
<211> 35
<212> DNA
<213> Artificial sequence
<220>
<223> Sequence of the junction region between Barnase and p3 in recombi
      nant fusion vector fd-3.
<220>
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<223> Sequence of the junction region between Barnase and p3 in recombi
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35

<210> 78

<400> 77

atcagactgc aggcggtgcg gccgcagaaa ctgtt

<211> 11

<212> PRT

<213> Artificial sequence <220> <223> Amino acid sequence about the junction of Barnase and p3 coding r egions of recombinant fusion vector fd-3. <400> 78 Ile Arg Leu Gln Ala Ala Ala Ala Glu Thr Val 1 5 10 <210> 79 <211> 4 <212> PRT <213> Artificial sequence <220> <223> Factor Xa protease cleavage sequence. <220> <221> MISC_FEATURE

<222> (1)..(1)

<223> X can be either Ile or Leu.

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<221> MISC_FEATURE
<222> (1)..(4)
<223> Factor Xa proteolytic cleavage site.
<400> 79
Xaa Glu Gly Arg
1
??
??
(continued...)
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(continued...)

Serial No. 09 710 4444 File No. 8034 1040 By. 4784) Applicant(s): 8 12 440 System and 8441 Title: \$28 12 410 System and 8441 The Following, DUE in the USPTO, was received by the PTO Mail Room on the date stamped hereon: Express Mail abel No. 843 140 System and the Mailing by Express Mail (37 CFR 1.10) Cent of Mailing by Express Mail (37 CFR 1.10) Express Mail abel No. 841 (37 CFR 1.10) Peristance of Mailing by Express Mail (38 CFR 1.10) Peristance of Mailing by Express Mail (38 CFR 1.10) Peristance of Mailing by Express Mail (38 CFR 1.10) Peristance of Mailing by Express Mail (38 CFR 1.10) Peristance of Mailing by Express Mail (38 CFR 1.10) Peristance of Mailing Mai	



·8039/1090

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of:

Riechmann, et al.

Serial No.:

09/710,444

Filed:

November 10, 2000

Entitled:

"Selection System"

Examiner:

B. Celsa

Group Art Unit:

1627

Conf. No.:

2736

CERTIFICATE OF MAILING UNDER 37 C.F.R. § 1.8a

I hereby certify that this correspondence (and any paper or fee referred to as being enclosed) is being deposited with the United States Post Office as First Class Mail on the date indicated below in an envelope addressed to: U.S. Patent & Trademark Office, Box: Sequence, P.O. Box 2327, Arlington, VA 22202.

Mary Wilson

Name of Person Mailing Paper

U.S. Patent and Trademark Office

Box: Sequence P.O. Box 2327

Arlington, VA 22202

TRANSMITTAL LETTER

Enclosed for filing the above-identified patent application, please find the following documents:

- 1. Amendment in Response to Notice to Comply dated February 27, 2002:
- 2. Copy of Notice to Comply;
- Paper Copy of the Sequence Listing (59 pgs); 3.
- 4. Computer Readable Copy of the Sequence Listing;
- Sequence Statement Under 37 C.F.R. § 1.821(f) and (g); 5.
- Petition for Four Month's Extension of Time; 6.
- 7. Check in the amount of \$720.00; and
- 8. Return Post Card.

The Commissioner for Patents is hereby authorized to charge any additional fees or credit any overpayment in the total fees to Deposit Account No. 16-0085, Reference 8039/1090. A duplicate of this transmittal letter is enclosed for this purpose. Mak J. FitzGeroll

Respectfully submitted,

Rg. No. 45, 928 fi-

Kathleen Williams

Date: July **25**, 2002

> Docket# Response due

Statutory period

Palmer & Dodge LLP Patent Department

Name: Kathleen Williams Registration No.: 34,380 Customer No.: 29933

Palmer & Dodge LLP 111 Huntington Avenue

Boston, MA 02199-7613 Tel: 617-239-0100



8039/1090

PATENT

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Riechmann, et al.

Serial No.:

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November 10, 2000

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Examiner:

B. Celsa

Group Art Unit:

1627

Conf. No.:

2736

CERTIFICATE OF MAILING UNDER 37 C.F.R. § 1.82

I hereby certify that this correspondence (and any paper or fee referred to as being enclosed) is being deposited with the United States Post Office as First Class Mail on the date indicated below in an envelope addressed to: U.S. Patent & Trademark Office, Box: Sequence, P.O. Box 2327, Arlington, VA 22202.

Mary Wilson

Name of Person Mailing Paper

U.S. Patent and Trademark Office

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Arlington, VA 22202

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- Sequence Statement Under 37 C.F.R. § 1.821(f) and (g); 5.
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Respectfully submitted.

Rg. No. 45, 928 for

Date: July 25, 2002

Kathleen Williams

Name: Kathleen Williams Registration No.: 34,380 Customer No.: 29933 Palmer & Dodge LLP 111 Huntington Avenue

Boston, MA 02199-7613 Tel: 617-239-0100



8039/1090



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

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Riechmann, et al.

Serial No.:

09/710,444

Filed:

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Mary Wilson

Name of Person Mailing Paper

U.S. Patent and Trademark Office

Box: Sequence P.O. Box 2327

Arlington, VA 22202

PETITION FOR EXTENSION OF TIME

Dear Sir:

Applicant respectfully petitions under 37 C.F.R. § 1.136(a) for an Extension of Time of four months to file a Response in the above-identified patent application. This will serve to extend the time for filing the Response to the Notice to Comply mailed February 27, 2002 from March 27, 2002, up to and including July 27, 2002.

Pursuant to 37 C.F.R. § 1.17, enclosed is the requisite extension fee of \$720.00 for maintaining the pendency of the present application. The Commissioner of Patents is hereby authorized to charge any additional fees or credit any overpayment to Deposit Account No. 16-0085, Reference No. 8039/1090. A duplicate of this letter is enclosed for that purpose.

Respectfully submitted,

Make J. Fitz Gentl Ry. No. 45,928 for Kathler M. William

Date: July 25, 2002

Name: Kathleen Williams

Registration No.: 34,380 Customer No.: 29933

Palmer & Dodge LLP 111 Huntington Avenue Boston, MA 02199-7613

Tel: 617-239-0100



8039/1090

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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of:

Riechmann, et al.

Serial No.:

09/710,444

Filed:

November 10, 2000

Entitled:

"Selection System"

Examiner:

B. Celsa

Group Art Unit:

1627

Conf. No.:

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Mary Wilson

Name of Person Mailing Paper

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Respectfully submitted,

Mark J. FifzGenll Ry. No. 45,928 for Kathlen M. William

Date:

July 25, 2002

Name: Kathleen William

Name: Kathleen Williams Registration No.: 34,380 Customer No.: 29933 Palmer & Dodge LLP

111 Huntington Avenue Boston, MA 02199-7613

Tel: 617-239-0100



8039/1090

PATENT

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Application of: Riechmann, et al.

Serial No.:

09/710,444

Filed:

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Entitled:

"Selection System"

Examiner:

B. Celsa

Group Art Unit:

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Conf. No.:

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Mary Wilson

Name of Person Mailing Paper

Signature of Person Mailing Paper

U.S. Patent and Trademark Office

Box: Sequence P.O. Box 2327

Arlington, VA 22202

STATEMENT UNDER 37 C.F.R. §1.821 (f) and (g)

Sir:

This paper is submitted in response to the Notice to Comply mailed by the USPTO on February 27, 2002.

In accordance with 37 C.F.R. §1.821 (f) I hereby state that the paper copy and the computer readable form of the Sequence Listing submitted herewith in the above-identified patent application are supported in the application and contain no new matter. Thereby state that the information recorded in computer readable form is identical to the written sequence listing.

In accordance with 37 C.F.R. §1.821 (g), I hereby state that the computer readable form of the Sequence Listing submitted herewith contains no new matter. Mark J. Fitz Grall

Reg. No. 45, 928 for Kathley Williams

7/25/02

Kathleen Williams

Reg. No. 34,380

Attorney for Applicant

Palmer & Dodge LLP

111 Huntington Avenue

Boston, MA 02199-7613 Tel: (617) 239-0451

Fax: (617) 227-4420



8039/1090

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of:

Riechmann, et al.

Serial No.:

09/710,444

Filed:

November 10, 2000

Entitled:

"Selection System"

Examiner:

B. Celsa

Group Art Unit:

1627

Conf. No.:

2736

CERTIFICATE OF MAILING UNDER 37 C.F.R. § 1.8a

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Mary Wilson

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Arlington, VA 22202

AMENDMENT

Sir:

This is filed in response to the Examiner's Notice to Comply with nucleotide or amino acid sequence listing requirements mailed February 27, 2002 in the above noted U.S. Patent Application. Kindly enter the following amendments and remarks.

In the Specification:

Replace the paragraph at lines 14 to 23 on page 15 with the following replacement paragraph:

A sequence (PAGLSEGSTIEGRGAHE; SEQ ID NO: 1) comprising several proteolytic sites is inserted in the flexible glycine-rich region between the D2 and D3 domains of the phage p3. Incubation of the phage (fd-K108) under native conditions with trypsin, thermolysin or subtilisin now resulted in almost complete loss of infectivity (from 10^7 to < 10 TU/ml) and incubation with Glu-C and chymotrypsin resulted in a major loss (from 10^7 to 10^4 TU/ml). This indicates that these proteases cleave the new linker. However incubation with Factor Xa, Arg-C or thrombin did not lead to a loss in infectivity, despite the presence of potential cleavage sites

for these enzymes. Presumably the presence of the D2 and D3 domains may block access or cleavage for these enzymes in the case of the present polypeptide.

Replace Table 4, on page 24, with the following replacement Table 4:



TECH CENTER 1600/2900

Table 4

Table 4. Primer sequences

pklinker	5'GGCACCCTCAGAACGGTACCCCACCCTCAGAGGCCGGCTGGG CCGCCACCCTCAGAG 3' (SEQ ID NO: 2)
polyXafor	5'GGTGGCGGCCCAGCCGGCCTTTCTGAGGGGTCGACTATAGAA GGACGAGGGCCCAGCGAAGGAGGTGGGGTACCCCCTTCTGAGG GTGG 3' (SEQ ID NO: 3)
polyXaback	5'CCACCCTCAGAAGGGGGTACCCCACCTCCTTCGCTGGGCCCT CGTCCTTCTATAGTCGACCCCTCAGAAAGGCCGGCTGGGCCGC CACC 3' (SEQ ID NO: 4)
fdPCRBack	5'GCGATGGTTGTCATTGTCGGC 3' (SEQ ID NO: 5)
LIBSEQfor	5'AAAAGAAACGCAAAGACACCACGG 3' SEQ ID NO: 6)
LIBSEQback	5'CCTCCTGAGTACGGTGATACACC 3' (SEQ ID NO: 7)
LSPAf or	5'GTAAATTCAGAGACTGCGCTTTCC 3' (SEQ ID NO: 8)
LSPAback	5'ATTTTCGGTCATAGCCCCCTTATTAG 3' (SEQ ID NO: 9)
Flagprimer	5'CAACGGGCGGCCGCAGACTACAAGGATGACGACGACAAGG AAACTGTTGAAAGTTGTTTAGCAA 3' (SEQ ID NO: 10)
RECGLYfor	5'CCCCTCAGAAAGGCCGGCTGGGCCGCCGCCAGCATTGACAG GAGGTTCAGG 3' (SEQ ID NO: 11)
RECGLYback	5'GAAGGAGGTGGGTACCCGGTTCCGAGGGTGGTTCCGGTTC CGGTGATTTTG 3' (SEQ ID NO: 12)
delcKpn	5'CCCTCGGAACCGGTACCCCAGCTGCTTCGTGGGCCC 3' (SEQ ID NO: 13)
Barnasefor	5'CTGGCGGCCCAGCCGGCCCTGCACAGGTTATCAACACG TTTGAC 3' (SEQ ID NO: 14)
BarnaseH102Aba	5'CTCGGAACCGGTACCTCTGATTTTTGTAAAGGTCTGATAAGC G 3' (SEQ ID NO: 15)
ck	
villinfor	5'GGCGGCCCAGCCGGCCTTTCTCTCTCTGACGAGGACTTCAAG GC 3' (SEQ ID NO: 16)
villinback	5'CCTCGGAACCGGTACCGAAGAGTCCTTTCTCCTTCTTGAGG 3' (SEQ ID NO: 17)

-Replace the paragraph at page 30, lines 4-14 with the following replacement paragraph:

- --1: TACGCCAAGCTTGCATGC (SEQ ID NO: 18);
- 2: CTGCACCTGGGCCATGG (SEQ ID NO: 19);
- 3: GATTACGCCAAGCTTTG (SEQ ID NO: 20);
- 4: GATTACGCC*AAGCTT*GCATGCANNDDCTNTDTCAAGGAGACAGTCATAATGARRN NBCTATTGSYAAYRSYASYASYAGBNTTGTTATTACTCSYANYCVNNCYGD*CCATGG* CCCAGGTGCAGCTG (SEQ ID NO: 21);
- 5: GATTACGCC*AAGCTT*TGNNNNCTTTTTTWWGGAGATTTCAACRTGARAARATTAT TATTCSYAATTSYTTAGTTSYTSYTTTCTWTGYGGYCCAGCCGG*CCATGG*CCCAGGT GCA. (SEQ ID NO: 22)
- 6: CTTTATGCTTCCGGCTCG. (SEQ ID NO: 23)
- 7: CGGCCCCATTCAGATCC. (SEQ ID NO: 24)--

Replace Table 7, on pages 35 and 36, with the following replacement Table 7. Because portions of the original text are underlined, the present amendments are indicated by <u>double underlines</u>.

Table 7. Randomised and selected sequences.

The randomised DNA sequence is given from 5' to 3'; above and below it, the bases that differ from the given sequence in the signal sequences pe1B, 17, 19, 110 and 112 are indicated. The Shine-Delgarno sequence, the start codon and the last codon of the signal sequence, GCC, have been underlined. The HindIII and the NcoI restriction sites are in italics. The corresponding amino acid sequences are given below. Library I is initially designed from the pelB leader and library II from the g3 leader.

III-A. From library I

pelB (SEQ ID NO: 25) AATT A T AATAC

5' AAGCTTGCATGCANNDDCTNT DTCAAGGAGACAGTCATAAATGARRNNB CT (SEQ ID NO:26)

 17 (SEQ ID NO: 27)
 GCAT
 C
 G
 AGACG

 110 (SEQ ID NO: 28)
 CGGG
 G
 T
 GAGGG

 112 (SEQ ID NO: 29)
 CCAG
 C
 T
 GGCGG

pelB CCT CGGC GCCGCT GA GCGGC CAG C G (SEQ ID NO: 30)

ATTGSYAAYRSYASYAGBNTTGTTATTACTC SYANY CVNNCYGD*CCATGG*

<u>CC</u> 3' (SEQ ID NO: 31)

17 GC TGGT CT GT GA CC CC GGT C T (SEQ ID

NO: 32)

110 GC TGCT GT GC GG CC AT GCG C G (SEQ ID

NO: 33)

112 GT TAGC CG CT

GG CT GC CCC C

)

A (SEQ ID

NO: 34)

pelB MKYLLPTAAAGLLLLAAQPAMA (SEQ ID NO: 35)

17 KT AMVLVG PPGPS (SEQ ID NO: 36)

110 RG AMLVAG PIAPA (SEQ ID NO: 37)

112 RR VIAAVG LAPPT (SEQ ID NO: 38)

III-B. From library II

g3leader

GAGC '

TT

G A A (SEQ ID NO: 39)

5' AAGTTGNNNNCTTTTTT<u>WWGGAG</u>ATTTTCAAC<u>RTG</u>ARAARATTATTAT (SEQ ID NO: 40)

19

GGGC

TA

A GG (SEQ ID NO: 41)

GC CC GT CC A C C (SEQ ID NO: 42)

TCSYAATTSYTTAGTTSYTSYTTTCTWTGYGGYCCAGCCGGCCATGG CC3 $^{\circ}$ (SEQ ID NO: 43)

19 CT CC GT GC A T T (SEQ ID NO: 44)

g3 leader MKKLLFAIPLVVPF

YAAQPAMA (SEQ ID NO: 45)

19 RRLPVA

Y V V (SEQ ID NO: 46)

Replace the paragraph at lines 8-17 on page 38 with the following replacement paragraph:

The phage displaying the Stoffel fragment are incubated with primer 13 [TTT CGC AAG ATG TGG CGT] (SEQ ID NO: 47) comprising a 5' maleimidyl group and a 3' biotinylated nucleotide. After incubation the phage are captured on streptavidin-coated beads, with a yield of about 1-5% of infectious phage. This shows that primer can be chemically cross-linked to the phage, presumably via p8 protein as shown for the N-biotinoyl-N'(6-maleimidohexanoyl) bydrazide. The phage are then incubated with primer 1b [GCGAAGATGTGG] (SEQ ID NO: 48) comprising a 5' maleimidyl group in the presence of biotin-dUTP 2 and template 3 [AAA TAC AAC AAT AAA ACG CCA CAT CTT GCG] (SEQ ID NO: 49). Capture of the phage is dependent on presence of 1b, 2 and 3 (Table 8), but also on the inclusion of trypsin, which cleaves the helper phage to reduce non-specific phage isolation.

Replace the paragraph at page 39, lines 19-27 with the following replacement paragraph:

For the cloning of (poly)-peptide encoding DNA fragments and their display for selection between barnase and p3, the phage fd-3 is constructed (Fig. 5). Phage fd-3 comprises the H1021A mutant of barnase N-terminally fused to the p3 gene of phage fd.TET. Between the codon for the last residue of barnase and the first residue of p3 is the nucleotide sequence *CTG GAG* GCG GTG CGG CCG CA (SEQ ID NO: 50). This sequence contains a PstI DNA restriction site (in italics) for insertion of DNA fragments flanked by PstI restriction sites. The sequence further introduces a frame shift between barnase and p3, which prevents expression of the correct p3 reading frame in fd-3. Phage particles of phage fd-3 therefore do not display the infection protein p3 and are non-infectious.

Replace the paragraph at page 40, lines 8-23 with the following replacement paragraph:

Genomic DNA from the E. coli strain TG1 is amplified in 30 cycles of a polymerase chain reaction (PCR) with an annealing temperature of 48°C using the oligonucleotide SN6MIX (5'-GAG CCT GCA GAG CTC AGG NNN NNN-3'; SEQ ID NO: 51), which comprises 6 degenerate positions at the extendible 3' end to ensure random priming. In a second step of 30 PCR cycles with an annealing temperature of 52°C primary PCR products are extended by reamplification with the oligonucleotide XTND (5'-CGT GCG AGC CTG CAG AGC TCA GG-3'; SEQ ID NO: 52). Products with a length of around 150 bp from this reaction are purified from an agarose gel and reamplified in 30 PCR cycles using an annealing temperature of 52°C and the oligonucleotide XTND. These reamplified 150 bp fragments are partially digested with SacI (site indicated in bold in the oligonucleotides) and ligated for dimerisation. Ligated products are reamplified in a further 10 PCR cycles with an annealing temperature of 44°C followed by a 30 PCR cycles with an annealing temperature of 55°C using the oligonucleotide The annealing temperatures are chosen to discriminate against priming of the oligonucleotide in the middle of the dimerised fragments. The reaction product is size purified twice on an agarose gel to remove monomers and oligomers (non-dimers).

Replace the table on page 44 (Table 9) with the following replacement table:

Phage	Proteolytic	Barstar	bindg	Amino acid sequence
clone	selection	-DTT	+DTT	of inserts
TA-1.2	1xTr	yes	no	LQSSGDCVIS DTCIAGMAEA AACEEKFSSQ NVGLTITVTP CLSSA (SEQ ID NO: 53)
TA-2.25	2xTr	yes	no	LQSSGCGSSG SSINCLPCGA TSRGTSPLAS GLPSSATIHC LSSA (SEQ ID NO: 54)
TA-2.26	2xTr	yes	no	LQSSGDSAGC KNMTGGRLYA HTLEAIIPGF AVSAPACEPA (SEQ ID NO: 55)
TA-2.27	2xTr	yes	yes	LQSSGCVRLK RTSVNHQPDA WPEPHLKAAC EPA (SEQ ID NO: 56)
TA-2.30	2xTr	yes	no	LQSSGCGSSG SSINCLPCGA TSRGTSPLAS GLPSSATVQC LSSA (SEQ ID NO: 57)
TB-1.10	1xTh	yes	yes	LQSSGKIVQA GANIQDGCIM HGYCDTDTIV GENGHIGLSS A (SEQ ID NO: 58)
TB-1.11	1xTh	yes	yes	no insert, Barnase & p3 in frame
TB-2.33	2xTh	yes	no	LQSSGVCVIS DTCIAGTAEA AACEEKFSSQ NVGHTITETP CLSSA (SEQ ID NO: 59)
TB-2.34	2xTh	yes	no	LQSSGCGSSG SSINCLPCGA TSRGTSPLAS GLPSSATIQC LSSA (SEQ ID NO: 60)
TE-2.35	2xTh	yes	no	LQSSGQDSQR EHASHTAEDD CEDQTRIHQH IREVDFVDTP QEVDDCRAAL SSA (SEQ ID
NO: 61)				
TB-2.37	2xTh	yes	no	LQSSGCVRLK RTSVNHQPDA WPEPHLKAAC EPA (SEQ ID NO: 62)
TB-2.38	2xTh	yes	yes	LQSSGVRPA (SEQ ID NO: 63)
TB-2.39	2xTh	yes	no	LQSSGCGSS GSSINCLPCGA TSRGTSPLAS GLPSSATIQ CLSSA (SEQ ID NO: 64)

Replace the table at lines 12-29 on page 46 with the following replacement table:

Phage	Proteolytic	Barstarbindg	Amino acid sequence		
clone	selection	+DTT	of inserts		
B2-13 (SEQ ID NO	2xTr/Th O: 65)	yes	LQSSGTEVDR GNQQHDTNDR DFTHTPLSS A		
B2-14	2xTr/Th	yes	LQSSG5VAQG SSASVDVTAT NAVLSADSL SLGGGEPA (SEQ ID NO: 66)		
B2-22	2xTr/Th	yes	LQSSGGAVAV TPGPVLSSA (SEQ ID NO: 67)		
B2-23	2xTr/Th	yes	LQSSGHCRGK PVLCTHTA (SEQ ID NO: 68)		
B2-15	2xTr/Th	yes	LQSSGVRPA (SEQ ID NO: 69)		
B2-17	2xTr/Th	yes	no insert, Barnase & p3 in frame		
B2-20,21	2xTr/Th	yes	no insert, Barnase & p3 in frame		
B2-16,24	2xTr/Th	yes	LQSSGEPAPA HEAKPTEAPV AKAEAKPETP AHLSSA (SEQ ID NO: 70)		
B2-18	2xTr/Th	no	LQSSGCVRLK RTSVNHQPDA WPEPHLKAAC EPA (SEQ ID NO: 71)		
B2-19	2xTr/Th	no	LQSSGVVDWA KMREIADSIG AYLFVDMAHV AALSSA (SEQ ID NO: 72)		

Replace the paragraph at page 47, lines 8-10 with the following replacement paragraph:

Figure 2. The phagemid vectors pK1 and pK2. These vectors contain a protease cleavable sequence between D2 and D3 of the phage p3 protein. In pK1, D2 + D3 are in frame; in pK2, D3 is out of frame. Nucleotide and amino acid sequence for the polylinker regions are shown for pK1 (SEQ ID NO: 73 and SEQ ID NO: 74, respectively) and pK2 (SEQ ID NO: 75 and SEQ ID NO: 76, respectively).

Replace the paragraph at page 47, lines 21-23 with the following replacement paragraph:

Figure 5. The fd vector fd-3. The gene for the H102A mutant of Barnase is introduced by subcloning into fd-DOG [43] after PCR amplification with suitable oligonucleotides using the restriction sites ApaLI (at the Barnase 5' end) and NotI to create fd-3. The nucleotide and amino

acid sequence of the junction between Barnase and p3 sequences is shown in expanded view (SEQ ID NO 77 and SEQ ID NO: 78, respectively)

REMARKS

The amendments directed herein are made in order to add SEQ ID NOs corresponding to the SEQ ID NOs in the accompanying Sequence Listing. The amendments add no new matter.

Respectfully submitted: Rg. No. 45,928 for Kathleen M. William,

Mark J. Fitz Gentl

Kathleen M. Williams

Reg. No. 34, 380

Attorney for Applicant

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Version of amendments marked to show changes:

- Replace the paragraph at lines 14 to 23 on page 15 with the following replacement paragraph:

-- A sequence (PAGLSEGSTIEGRGAHE; SEQ ID NO: 1) comprising several proteolytic sites is inserted in the flexible glycine-rich region between the D2 and D3 domains of the phage p3. Incubation of the phage (fd-K108) under native conditions with trypsin, thermolysin or subtilisin now resulted in almost complete loss of infectivity (from 10⁷ to < 10 TU/ml) and incubation with Glu-C and chymotrypsin resulted in a major loss (from 10⁷ to 10⁴ TU/ml). This indicates that these proteases cleave the new linker. However incubation with Factor Xa, Arg-C or thrombin did not lead to a loss in infectivity, despite the presence of potential cleavage sites for these enzymes. Presumably the presence of the D2 and D3 domains may block access or cleavage for these enzymes in the case of the present polypeptide.--

- Replace Table 4, on page 24, with the following replacement Table 4:

--Table 4

Table 4. Primer sequences

pklinker	5'GGCACCCTCAGAACGGTACCCCACCCTCAGAGGCCGGCTGGG CCGCCACCCTCAGAG 3' (SEQ ID NO: 2)
polyXafor	5'GGTGGCGGCCCAGCCGGCCTTTCTGAGGGGTCGACTATAGAA GGACGAGGGCCCAGCGAAGGAGGTGGGGTACCCCCTTCTGAGG GTGG 3' (SEQ ID NO: 3)
polyXaback	5'CCACCCTCAGAAGGGGGTACCCCACCTCCTTCGCTGGGCCCT CGTCCTTCTATAGTCGACCCCTCAGAAAGGCCGGCTGGGCCGC CACC 3' (SEQ ID NO: 4)
fdPCRBack	5'GCGATGGTTGTCATTGTCGGC 3' (SEQ ID NO: 5)
LIBSEQfor	5'AAAAGAAACGCAAAGACACCACGG 3' (SEQ ID NO: 6)
LIBSEQback	5'CCTCCTGAGTACGGTGATACACC 3' (SEQ ID NO: 7)
LSPAf or	5'GTAAATTCAGAGACTGCGCTTTCC 3' (SEQ ID NO: 8)
LSPAback	5'ATTTTCGGTCATAGCCCCCTTATTAG 3' (SEQ ID NO: 9)
Flagprimer	5'CAACGGGCGGCCGCAGACTACAAGGATGACGACGACAAGG AAACTGTTGAAAGTTGTTTAGCAA 3' (SEQ ID NO: 10)
RECGLYfor	5'CCCCTCAGAAAGGCCGGCTGGGCCGCCGCCAGCATTGACAG GAGGTTCAGG 3' (SEQ ID NO: 11)
RECGLYback	5'GAAGGAGGTGGGTACCCGGTTCCGAGGGTGGTTCCGGTTC CGGTGATTTTG 3' (SEQ ID NO: 12)
delcKpn	5'CCCTCGGAACCGGTACCCCAGCTGCTTCGTGGGCCC 3' (SEQ ID NO: 13)
Barnasefor	5'CTGGCGGCCCAGCCGGCCCTGCACAGGTTATCAACACG TTTGAC 3' (SEQ ID NO: 14)
BarnaseH102Aback	5'CTCGGAACCGGTACCTCTGATTTTTGTAAAGGTCTGATAAGC G 3' (SEQ ID NO: 15)
villinfor	5'GGCGGCCCAGCCGGCCTTTCTCTCTCTGACGAGGACTTCAAGGCC3' (SEQ ID NO: 16)
villinback	5'CCTCGGAACCGGTACCGAAGAGTCCTTTCTCCTTCTTGAGG 3' (SEQ ID NO: 17)

- -Replace the paragraph at page 30, lines 4-14 with the following replacement paragraph:
- --1: TACGCCAAGCTTGCATGC (SEQ ID NO: 18);
- 2: CTGCACCTGGGCCATGG (SEQ ID NO: 19);
- 3: GATTACGCCAAGCTTTG (SEQ ID NO: 20);
- 4: GATTACGCC*AAGCTT*GCATGCANNDDCTNTDTCAAGGAGACAGTCATAATGARRN NBCTATTGSYAAYRSYASYASYAGBNTTGTTATTACTCSYANYCVNNCYGD*CCATGG* CCCAGGTGCAGCTG (SEQ ID NO: 21);
- 5: GATTACGCC*AAGCTT*TGNNNNCTTTTTTWWGGAGATTTCAACRTGARAARATTAT TATTCSYAATTSYTTAGTTSYTSYTTTCTWTGYGGYCCAGCCGG*CCATGG*CCCAGGT GCA. (SEQ ID NO: 22)
- 6: CTTTATGCTTCCGGCTCG. (SEQ ID NO: 23)
- 7: CGGCCCCATTCAGATCC. (SEQ ID NO: 24)--
- Replace Table 7, on pages 35 and 36, with the following replacement Table 7. Because portions of the original text are underlined, the present amendments are indicated by <u>double underlining</u>.

-Table 7. Randomised and selected sequences.

The randomised DNA sequence is given from 5' to 3'; above and below it, the bases that differ from the given sequence in the signal sequences pe1B, 17, 19, 110 and 112 are indicated. The Shine-Delgarno sequence, the start codon and the last codon of the signal sequence, GCC, have been underlined. The HindIII and the NcoI restriction sites are in italics. The corresponding amino acid sequences are given below. Library I is initially designed from the pelB leader and library II from the g3 leader.

III-A. From library I

III-A. Fro	m library .	i						
pelB (SEQ	ID NO: 25) AATT	A			T		AATAC
5' AAGO	TTGCAT	GCANN	NDDCTN7	T DTC	AAGGAGA	CAGTCATA	AA <u>ATG</u> A	RRNNB CT
(SEQ ID N	<u>O:26)</u>				•	,		
·	···							
17 <u>(SEQ II</u>	NO: 27)	GCAT	C	G			AGACG	÷
110 (SEQ I	D NO: 28)	CGGC	G G	T			GAGGG	j
112 (SEQ I	D NO: 29)	CCAC	3 C	T			GGCGG	} ·
pelB CC	r cggc	GCCC	GCT GA		GCGGC	CAG C	G (SE	Q ID NO: 30)
ATTGS	YAAYRSY	ASYA	SYAGBN	TTGTT	TATTACTC	SYANY	CVNNC	YGD <i>CCATG<u>G</u></i>
CC 3' (SEQ ID NO: 31)								
17 GC	C TGGT	CT	GT		GA CC	CC GGT	C	T (SEQ ID
NO: 32)								
110 GO	C TGCT	GT	GC		GG CC	AT GCG	C	G (SEQ ID
NO: 33)								

112 GT TAGC CG CT

GG CT GC CCC C

A (SEQ ID

NO: 34)

pelB MKYLLPTAAAGLLLLAAQPAMA (SEQ ID NO: 35)

17 KT AMVLVG PPGPS (SEQIDNO: 36)

110 RG AMLVAG PIAPA (SEQ ID NO: 37)

112 RR VIAAVG LAPPT (SEQ ID NO: 38)

III-B. From library II

g3leader

GAGC

TT

G A A (SEQ ID NO: 39)

5' AAGTTGNNNNCTTTTTT<u>WWGGAG</u>ATTTTCAAC<u>RTG</u>ARAARATTATTAT

(SEQ ID NO: 40)

<u>19</u>

GGGC

TA

A G G (SEQ ID NO: 41)

GC CC GT CC A C C (SEQ ID NO: 42)

TCSYAATTSYTTTAGTTSYTSYTTTCTWTGYGGYCCAGCCGGCCATGG CC3'

(SEQ ID NO: 43)

19 CT CC GT GC A T T (SEQ ID NO: 44)

g3 leader MKKLLFAIPLVVPF

YAAQPAMA (SEQ ID NO: 45)

19

RR LP VA

Y V V (SEQ ID NO: 46)

--Replace the paragraph at lines 8-17 on page 38 with the following replacement paragraph. Please note that the sequences were presented in brackets in the original filed application. The brackets are replaced by parenthesis herein.

--The phage displaying the Stoffel fragment are incubated with primer 13 (TTT CGC AAG ATG TGG CGT) (SEQ ID NO: 47) comprising a 5' maleimidyl group and a 3' biotinylated nucleotide. After incubation the phage are captured on streptavidin-coated beads, with a yield of about 1-5% of infectious phage. This shows that primer can be chemically cross-linked to the phage, presumably via p8 protein as shown for the N-biotinoyl-N'(6-maleimidohexanoyl) bydrazide. The phage are then incubated with primer 1b (GCGAAGATGTGG) (SEQ ID NO: 48) comprising a 5' maleimidyl group in the presence of biotin-dUTP 2 and template 3 (AAA TAC AAC AAT AAA ACG CCA CAT CTT GCG) (SEQ ID NO: 49). Capture of the phage is dependent on presence of 1b, 2 and 3 (Table 8), but also on the inclusion of trypsin, which cleaves the helper phage to reduce non-specific phage isolation.--

- Replace the paragraph at page 39, lines 19-27 with the following replacement paragraph:

--For the cloning of (poly)-peptide encoding DNA fragments and their display for selection between barnase and p3, the phage fd-3 is constructed (Fig. 5). Phage fd-3 comprises the H1021A mutant of barnase N-terminally fused to the p3 gene of phage fd.TET. Between the codon for the last residue of barnase and the first residue of p3 is the nucleotide sequence CTG GAG GCG GTG CGG CCG CA (SEQ ID NO: 50). This sequence contains a PstI DNA restriction site (in italics) for insertion of DNA fragments flanked by PstI restriction sites. The sequence further introduces a frame shift between barnase and p3, which prevents expression of the correct p3 reading frame in fd-3. Phage particles of phage fd-3 therefore do not display the infection protein p3 and are non-infectious.--

- Replace the paragraph at page 40, lines 8-23 with the following replacement paragraph:

--Genomic DNA from the E. coli strain TG1 is amplified in 30 cycles of a polymerase chain reaction (PCR) with an annealing temperature of 48°C using the oligonucleotide SN6MIX (5'-GAG CCT GCA GAG CTC AGG NNN NNN-3'; SEQ ID NO: 51), which comprises 6 degenerate positions at the extendible 3' end to ensure random priming. In a second step of 30 PCR cycles with an annealing temperature of 52°C primary PCR products are extended by reamplification with the oligonucleotide XTND (5'-CGT GCG AGC CTG CAG AGC TCA GG-3'; SEQ ID NO: 52). Products with a length of around 150 bp from this reaction are purified from an agarose gel and reamplified in 30 PCR cycles using an annealing temperature of 52°C and the oligonucleotide XTND. These reamplified 150 bp fragments are partially digested with SacI (site indicated in bold in the oligonucleotides) and ligated for dimerisation. Ligated products are reamplified in a further 10 PCR cycles with an annealing temperature of 44°C followed by a 30 PCR cycles with an annealing temperature of 55°C using the oligonucleotide The annealing temperatures are chosen to discriminate against priming of the XTND. oligonucleotide in the middle of the dimerised fragments. The reaction product is size purified twice on an agarose gel to remove monomers and oligomers (non-dimers).--

- Replace the table on page 44 (Table 9) with the following replacement table:

Phage	Proteolytic	Barstarl	bindg	Amino acid sequence
clone	selection	-DTT	+DTT	of inserts
TA-1.2	1xTr	yes	no	LQSSGDCVIS DTCIAGMAEA AACEEKFSSQ NVGLTITVTP CLSSA (SEQ ID NO: 53)
TA-2.25	2xTr	yes	no	LQSSGCGSSG SSINCLPCGA TSRGTSPLAS GLPSSATIHC LSSA (SEQ ID NO: 54)
TA-2.26	2xTr	yes	no	LQSSGDSAGC KNMTGGRLYA HTLEAIIPGF AVSAPACEPA (SEQ ID NO: 55)
TA-2.27	2xTr	yes	yes	LQSSGCVRLK RTSVNHQPDA WPEPHLKAAC EPA (SEQ ID NO: 56)
TA-2.30	2xTr	yes	no	LQSSGCGSSG SSINCLPCGA TSRGTSPLAS GLPSSATVQC LSSA (SEQ ID NO: 57)
TB-1.10	1xTh	yes	yes	LQSSGKIVQA GANIQDGCIM HGYCDTDTIV GENGHIGLSS A <u>(SEQ ID NO: 58)</u>
TB-1.11	1xTh	yes	yes	no insert, Barnase & p3 in frame
TB-2.33	2xTh	yes	no	LQSSGVCVIS DTCIAGTAEA AACEEKFSSQ NVGHTITETP CLSSA (SEQ ID NO: 59)
TB-2.34	2xTh	yes	no	LQSSGCGSSG SSINCLPCGA TSRGTSPLAS GLPSSATIQC LSSA (SEQ ID NO: 60)
TE-2.35	2xTh	yes	no	LQSSGQDSQR EHASHTAEDD CEDQTRIHQH IREVDFVDTP QEVDDCRAAL SSA (SEQ ID
NO: 61)				
TB-2.37	2xTh	yes	no	LQSSGCVRLK RTSVNHQPDA WPEPHLKAAC EPA <u>(SEQ ID NO: 62)</u>
TB-2.38	2xTh	yes	yes	LQSSGVRPA (SEQ ID NO: 63)
TB-2.39	2xTh	yes	no	LQSSGCGSS GSSINCLPCGA TSRGTSPLAS GLPSSATIQ CLSSA (SEQ ID NO: 64)

- Replace the table at lines 12-29 on page 46 with the following replacement table:

Phage clone	Proteolytic selection	Barstarbindg +DTT	Amino acid sequence of inserts
B2-13 (SEQ ID NO	2xTr/Th <u>D: 65)</u>	yes	LQSSGTEVDR GNQQHDTNDR DFTHTPLSS A
B2-14	2xTr/Th	yes	LQSSG5VAQG SSASVDVTAT NAVLSADSL SLGGGEPA (SEQ ID NO: 66)
B2-22	2xTr/Th	yes	LQSSGGAVAV TPGPVLSSA (SEQ ID NO: 67)
B2-23	2xTr/Th	yes	LQSSGHCRGK PVLCTHTA (SEQ ID NO: 68)
B2-15	2xTr/Th	yes	LQSSGVRPA (SEQ ID NO: 69)
B2-17	2xTr/Th	yes	no insert, Barnase & p3 in frame
B2-20,21	2xTr/Th	yes	no insert, Barnase & p3 in frame
B2-16,24	2xTr/Th	yes	LQSSGEPAPA HEAKPTEAPV AKAEAKPETP AHLSSA (SEQ ID NO: 70)
B2-18	2xTr/Th	no	LQSSGCVRLK RTSVNHQPDA WPEPHLKAAC EPA (SEQ ID NO: 71)
B2-19	2xTr/Th	no	LQSSGVVDWA KMREIADSIG AYLFVDMAHV AALSSA (SEQ ID NO: 72)

- Replace the paragraph at page 47, lines 8-10 with the following replacement paragraph:
- -- Figure 2. The phagemid vectors pK1 and pK2. These vectors contain a protease cleavable sequence between D2 and D3 of the phage p3 protein. In pK1, D2 + D3 are in frame; in pK2, D3 is out of frame. Nucleotide and amino acid sequence for the polylinker regions are shown for pK1 (SEQ ID NO: 73 and SEQ ID NO: 74, respectively) and pK2 (SEQ ID NO: 75 and SEQ ID NO: 76, respectively). --
- Replace the paragraph at page 47, lines 21-23 with the following replacement paragraph. Please note that number "43" was presented in brackets in the originally filed application. The brackets are replaced with parentheses herein.
- -- Figure 5. The fd vector fd-3. The gene for the H102A mutant of Barnase is introduced by subcloning into fd-DOG (43) after PCR amplification with suitable oligonucleotides using the

restriction sites ApaLI (at the Barnase 5' end) and NotI to create fd-3. The nucleotide and amino acid sequence of the junction between Barnase and p3 sequences is shown in expanded view (SEQ ID NO 77 and SEQ ID NO: 78, respectively).--



UNITED STATES PATENT AND TRADEMARK OFFICE

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER OF PATENTS AND TRADEMARKS Washington, D.C. 20231 www.uspto.gov

<u> </u>			* •	
APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/710,444	11/10/2000	Lutz Riechmann	8654/1090	5253
1	7590 02/27/2002			
Palmer & Dodge LLP		EXAMI	NER ()	
	e Beacon Street ston, MA 02109-3190		CELSA, BE	NNETT M
			ART UNIT	PAPER NUMBER
			1627	
			DATE MAILED: 02/27/2002	

Please find below and/or attached an Office communication concerning this application or proceeding.

way

Palmer & Dodge LLP Drop
Patent Department

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Palmer & Dødge, LLP Petent Department



UNITED STATES DEPARTMENT OF COMMERCE Patent and Trademark Office COMMISSIONER OF PATENTS AND TRADEMARKS Washington, D.C. 20231

SERIAL NUMBER	FILING DATE	FIRST NAMED APPLICANT	ATTORNEY DOCKET NO.
09/ 710,444			
			FXAMINER

EXAMINER			
ART UNIT	PAPER NUMBER		
1627	5		

Please find below a communication from the EXAMINER in charge of this application

Sequence Rule Compliance: NOTICE TO COMPLY

This application fails to comply with the sequence rule quirements of 37 C.F.R. §§ 1.821-1.825 for the reason(s) set forth on the attached Notice To Comply With Requirements For Patent Applications Containing Nucleotide Sequence And/Or Amino Acid Sequence Disclosures. This application encompasses sequences needing sequence identifiers (e.g. see pages 9, 15, 24, 30, 35, 36, 38, figures etc.).

APPLICANT IS GIVEN 30 days FROM THE DATE OF THIS LETTER WITHIN WHICH TO COMPLY WITH THE SEQUENCE RULES, 37 C.R.F. §§ 1.821-1.825. Failure to comply with these requirements will result in ABANDONMENT of the application under 37 C.F.R. § 1.821(g). Extensions of time may be obtained by filing a petition accompanied by the extension fee under the provisions of 37 C.F.R. § 1.136. In no case may an applicant extend the period for response beyond the six month statutory period. Direct the response to the undersigned. Applicant is requested to return a copy of the attached Notice to Comply with the response.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Examiner Celsa whose telephone number is (703) 305-7556 If the examiner cannot be reached, inquiries can be directed to Supervisory Patent Examiner Venkat whose telephone number is (703) 308-0570. The fax number for the organization where this application or proceeding is assigned is (703) 308-4242. Any inquiry of a general nature or relating to the status of this application should be directed to the Group receptionist whose telephone number is (703) 308-0196.

Bennett Celsa (au 1627) (Feb. 25,2002)

BENNETT CELSA PRIMABY EXAMINER

MINER

Application No.09/710, 444

NOTICE TO COMPLY WITH REQUIREMENTS FOR PATENT APPLICATIONS CONTAINING NUCLEOTIDE SEQUENCE AND/OR AMINO ACID SEQUENCE DISCLOSURES

The nucleotide and/or amino acid sequence disclosure contained in this application does not comply with the requirements for such a disclosure as set forth in 37 CFR 1.821 - 1.825 for the following reason(s):

2	or the following reason(s):	to requirements
M	•	
1. This application dearly fails	s to comply with the requirements of 37 CFR 1.821 -	
' directed to these regulations, p	oublished at 1114 OG 29, May 15, 1990 and at 55 FR	1.825. Applicant's attention is
2	29, May 15, 1990 and at 55 FR	R 18230, May 1, 1990.
== 2. This application does not co	intain, as a separate part of the disclosure on pages.	4
required by 37 CFR 1.821(c).	ontain, as a separate part of the disclosure on paper of	copy, a "Sequence Listing" as
· 1 () 1		
3. A copy of the "Sequence Lies	ting" in computer readable form has not been submitt	
Tanadactice Its	ung in computer readable form has not been submitt	ted as required by 27
4. A copy of the "Sequence List	ing" in computer readable form has been submitted.	· ·
computer readable form dode as	ing in computer readable form has been submitted.	However the contact - 44
attached marked-up copy of the	of comply with the requirements of 37 CFR 1.822 and Raw Sequence Listing."	Vor 1 823 as indicated
	. Train Sequence Listing."	as indicated on the
5 The second	·	•
. The computer readable form the	hat has been filed with this application has been for	
submitted as required to an the at	hat has been filed with this application has been foun	id to be damaged and/or
submitted as required by 37 CFR	that that been filed with this application has been foun tached CRF Diskette Problem Report. A substitute of 1.825(d).	computer readable form must be
<u> </u>		
6. The paper copy of the "Sequen	ice Liefings in a sur	· <u>.</u>
as required by 37 CFR 1.821(e)	ice Listing" is not the same as the computer readable	form of the "Sequence Listing"
102.(0).		orderice Listing
7. Other: ———		
. Outer:	·	
Applicant many		
Applicant must provide:		
		•
An initial or substitute computes an		·
The second secon	dable form (CRF) copy of the "Sequence Listing"	. •
An initial or substitute paper conver	the *Cook	
specification specification	the "Sequence Listing", as well as an amendment di	recting its entry into the
KTI		endy into the

A statement that the content of the paper and computer readable copies are the same and, where applicable, include no new matter, as required by 37 CFR 1.821(e) or 1.821(f) or 1.821(g) or 1.825(b) or 1.825(d) For questions regarding compliance with these requirements, please contact:

For Rules Interpretation, call (703) 308-1123 For CRF submission help, call (703) 308-4212 For Patentin software help, call (703) 308-6856

Please return a copy of this notice with your response.

Attachment for PTO-948 (Rev. 03/01, or earlier) 6/18/01

The below text replaces the pre-printed text under the heading, "Information on How to Effect Drawing Changes," on the back of the PTO-948 (Rev. 03/01, or earlier) form.

INFORMATION ON HOW TO EFFECT DRAWING CHANGES

1. Correction of Informalities - 37 CFR 1.85

New corrected drawings must be filed with the changes incorporated therein. Identifying indicia, if provided, should include the title of the invention, inventor's name, and application number, or docket number (if any) if an application number has not been assigned to the application. If this information is provided, it must be placed on the front of each sheet and centered within the top margin. If corrected drawings are required in a Notice of Allowability (PTOL-37), the new drawings MUST be filed within the THREE MONTH shortened statutory period set for reply in the Notice of Allowability. Extensions of time may NOT be obtained under the provisions of 37 CFR 1.136(a) or (b) for filing the corrected drawings after the mailing of a Notice of Allowability. The drawings should be filed as a separate paper with a transmittal letter addressed to the Official Draftsperson.

2. Corrections other than Informalities Noted by Draftsperson on form PTO-948.

All changes to the drawings, other than informalities noted by the Draftsperson, MUST be made in the same manner as above except that, normally, a highlighted (preferably red ink) sketch of the changes to be incorporated into the new drawings MUST be approved by the examiner before the application will be allowed. No changes will be permitted to be made, other than correction of informalities, unless the examiner has approved the proposed changes.

Timing of Corrections

Applicant is required to submit the drawing corrections within the time period set in the attached Office communication. See 37 CFR 1.85(a).

Failure to take corrective action within the set period will result in ABANDONMENT of the application.

SEQUENCE LISTING

- <110> Riechmann, Lutz
 Kristensen, Peter
 Jestin, Jean-Luc
 Winter, Gregory
- <120> Selection System
- <130> 8039/1090
 - <140> 09/710,444
 - <141> 2000-11-10
 - <150> GB 9810223.9
 - <151> 1998-05-13
 - <150> GB 9810228.8
 - <151> 1998-05-13
 - <150> PCT/GB99/01526
 - <151> 1999-05-13
 - <160> 78
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<213> Bacillus amyloliquefaciens

<400> 15

<210> 16

<211> 44

<212> DNA

<213> Gallus gallus

<400> 16

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<210> 17

<211> 41

<212> DNA

<213> Gallus gallus

<400> 17

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<211> 18

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<213> Artificial

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                                                                  17
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<213> Erwinia chrysanthemi
<220>
<221> misc_feature
<223> n at positions 23, 24, 29, 55, 56, 81, 97, 101, and 102 can be G,
       A, T or C
<220>
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<222> (23)..(23)
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<220>
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<221> misc_feature

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<223> n at position 24 can be G, A, T or C
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<221> misc_feature
<222> (29)..(29)
<223> n at position 29 can be G, A, T or C
<220>
<221> misc_feature
<222> (55)..(55)
<223> n at position 55 can be G, A, T or C
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<223> n at position 56 can be G, A, T or C
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<221> misc_feature

<222> (81)..(81)

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<221> misc_feature
<222> (97)..(97)
<223> n at position 97 can be G, A, T or C
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<222> (101)..(101)
<223> n at position 101 can be G, A, T or C
<220>
<221> misc_feature
<222> (102)..(102)
<223> n at position 102 can be G, A, T or C
<400> 21
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                                                                      60
ttgsyaayrs yasyasyagb nttgttatta ctcsyanycv nncygdccat ggcccaggtg
                                                                     120
cagctg
                                                                     126
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<210> 22

<211> 117

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<213> Bacteriophage M13mp18
<220>
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<223> Nucleotide at position 18 can be G, A, T or C.
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<221> misc_feature
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<220>
<221> misc_feature
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<223> Nucleotide at position 21 can be G, A, T or C.
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gattac	gcca agctttgnnn ncttttttww ggagattttc aacrtgaraa rattattatt	60
csyaat	tsyt ttagttsyts ytttctwtgy ggyccagccg gccatggccc aggtgca	117
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<213>	Artificial sequence	
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ctttat	gett eeggeteg	18
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7211/	17	
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<212>	DNA	
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<210> 25

<211> 50

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<210> 26

<211> 50

<212> DNA

<213> Artificial

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<220>
<221> misc_feature
<222> (15)..(15)
<223> n at position 15 can be G, A, T or C.
<220>
<221> misc_feature
<222> (20)..(20)
<223> n at position 20 can be G, A, T or C.
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<222> (46)..(46)
<223> n at position 46 can be G, A, T or C.
<400> 26
aagcttgcat gcannddctn tdtcaaggag acagtcataa tgarrnnbct
<210> 27
<211> 50
<212> DNA
<213> Artificial
<220>
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50

50

<213> Artificial

<210> 28

<211> 50

<212> DNA

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                                                                      50
<210> 29
<211> 50
<212> DNA
<213> Artificial
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                                                                      50
<210> 30
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<211> 55

<212> DNA

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<213> Artificial
<220>
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<223> n at position 22 can be G, A, T or C.

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<220>
<221> misc_feature
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                                                                     55
<210> 32
<211> 55
<212> DNA
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<213> Artificial

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<220>
<221> misc_feature
<222> (1)..(55)
<223> Randomized E. chrysanthemi pelB sequence
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                                                                    55
<210> 33
<211> 55
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<213> artificial
<220>
<221> misc_feature
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<223> Randomized E. chrysanthemi pelB sequence
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                                                                    55
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<210> 34

<211> 54

<212> DNA

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<223> n at position 43 can be G, A, T or C.
<220>
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<222> (44)..(44)
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<210> 35

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1 5 10 15

Ala Gln Pro Ala Met Ala

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<210> 36

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<210> 37

<211> 21

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10

15

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<210> 38

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10

15

Pro Thr Gln Pro Ala Met Ala

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<211> 50

<212> DNA

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<220>
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<222> (12)..(12)
<223> n at position 12 is can be G, A, t or C.
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<210> 41
<211> 50
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<220>

<212> DNA

<213> Artificial

. 50

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<221> misc_feature
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<223> Randomized bacteriophage M13 g3 sequence.

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aagctttggg gccttttttt aggagatttt caacatgaga agattattat

50

- <210> 42
- <211> 50
- <212> DNA
- <213> Artificial
- <220>
- <221> misc_feature
- <222> (1)..(50)
- <223> Randomized bacteriophage M13 g3 sequence
- <400> 42

tcgcaattcc tttagttgtt cctttctatg cggcccagcc ggccatggcc

- <210> 43
- <211> 50
- <212> DNA
- <213> Artificial

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<220>
<221> misc_feature
<222> (1)..(50)
<223> Randomized bacteriophage M13 g3 sequence
<400> 43
tcsyaattsy tttagttsyt sytttctwtg yggyccagcc ggccatggcc
                                                                     50
<210> 44
<211> 50
<212> DNA
<213> Artificial
<220>
<221> misc_feature
<222> (1)..(50)
<223> Randomized bacteriophage M13 g3 sequence
<400> 44
tcctaattcc tttagttgtt gctttctatg tggtccagcc ggccatggcc
                                                                     50
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<210> 45

<211> 22

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(22)

<223> Randomized bacteriophage M13 g3 sequence

<400> 45

Met Lys Lys Leu Leu Phe Ala Ile Pro Leu Val Val Pro Phe Tyr Ala

1

10

15

Ala Gln Pro Ala Met Ala

20

5

<210> 46

<211> 22

<212> PRT

<213> Artificial

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<222> (1)..(22)

<223> Randomized bacteriophage M13 g3 sequence

<400> 46

Met Arg Arg Leu Leu Ala Pro Pro Val Ala Val Pro Phe Tyr Val

1

10

15

Val Gln Pro Ala Met Ala

20

5

<210> 47

<211> 18

<212> DNA

<213> artificial

<220>

<221> misc_feature

<222> (1)..(18)

<223> Synthetic oligonucleotide primer used as substrate for Stoffel fr agment of Thermus aquaticus DNA polymerase I

<400> 47

tttcgcaaga tgtggcgt

```
<210> 48
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<211> 12

<212> DNA

<213> Artificial

<220>

<221> misc_feature

<222> (1)..(12)

<223> Synthetic primer used as substrate for Stoffel fragment of Thermu s aquaticus DNA polymerase I

<400> 48

gcgaagatgt gg

12

<210> 49

<211> 30

<212> DNA

<213> artificial

<220>

<221> misc_feature

<222> (1)..(30)

<223> Synthetic oligonucleotide primer used as substrate for Thermus aquaticus DNA polymerase I

<400> 49

aaatacaaca ataaaacgcc acatcttgcg

30

<210> 50

<211> 20

<212> DNA

<213> Artificial

<220>

<221> misc_feature

<222> (1)..(20)

<223> Synthetic oligonucleotide sequence insert containing PstI restric tion site and frame shift for H102A mutant barnase fusion construct fused to p3 gene of phage fd-3.

<400> 50

ctgcaggcgg tgcggccgca

20

<210> 51

<211> 24

<212> DNA

<213> artificial

<220>

<221> misc_feature

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<222> (1)..(24)
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<223> Synthetic oligonucleotide used for random priming

<220>

<221> misc_feature

<222> (19)..(19)

<223> n at position 19 can be G, A, T or C.

<220>

<221> misc_feature

<222> (20)..(20)

<223> n at position 20 can be G, A, T or C.

<220>

<221> misc_feature

<222> (21)..(21)

<223> n at position 21 can be G, A, T or C.

<220>

<221> misc feature

<222> (22)..(22)

<223> n at position 22 can be G, A, T or C.

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<220>
<221> misc_feature
<222> (23)..(23)
<223> n at position 23 can be G, A, T or C.
<220>
<221> misc_feature
<222> (24)..(24)
<223> n at position 24 can be G, A, T or C.
<400> 51
gagcctgcag agctcaggnn nnnn
                                                                    24
<210> 52
<211> 23
<212> DNA
<213> artificial
<220>
<221> misc_feature
<222> (1)..(23)
```

<223> Synthetic PCR primer used to re-amplify randomly amplified E. col

i genomic DNA sequences.

<400> 52

cgtgcgagcc tgcagagctc agg

23

<210> 53

<211> 45

<212> PRT

<213> artificial

<220>

<221> MISC_FEATURE

<222> (1)..(45)

<223> Barstar binding barnase-p3 fusion insert

<400> 53

Leu Gln Ser Ser Gly Asp Cys Val Ile Ser Asp Thr Cys Ile Ala Gly

1 5 10 15

Met Ala Glu Ala Ala Cys Glu Glu Lys Phe Ser Ser Gln Asn Val
20 25 30

Gly Leu Thr Ile Thr Val Thr Pro Cys Leu Ser Ser Ala

35 40 45

<210> 54

<211> 44

<212> PRT

<213> artificial

<220>

<221> MISC_FEATURE

<222> (1)..(44)

<223> Barstar binding barnase-p3 fusion insert

<400> 54

1

Leu Gln Ser Ser Gly Cys Gly Ser Ser Gly Ser Ser Ile Asn Cys Leu

10

15

Pro Cys Gly Ala Thr Ser Arg Gly Thr Ser Pro Leu Ala Ser Gly Leu

20

5

25

30

Pro Ser Ser Ala Thr Ile His Cys Leu Ser Ser Ala

35

40

<210> 55

<211> 40

<212> PRT

<213> artificial

<220>

<221> MISC_FEATURE

<222> (1)..(40)

<223> Barstar binding barnase-p3 fusion insert

<400> 55

Leu Gln Ser Ser Gly Asp Ser Ala Gly Cys Lys Asn Met Thr Gly Gly

1 5 10 15

Arg Leu Tyr Ala His Thr Leu Glu Ala Ile Ile Pro Gly Phe Ala Val 20 25 30

Ser Ala Pro Ala Cys Glu Pro Ala

35 40

<210> 56

<211> 33

<212> PRT

<213> artificial

<220>

<221> MISC_FEATURE

<222> (1)..(33)

<223> Barstar binding barnase-p3 fusion insert

<400> 56

Leu Gln Ser Ser Gly Cys Val Arg Leu Lys Arg Thr Ser Val Asn His

1

10

15

Gln Pro Asp Ala Trp Pro Glu Pro His Leu Lys Ala Ala Cys Glu Pro

20

25

30

Ala

<210> 57

<211> 44

<212> PRT

<213> artificial

<220>

<221> MISC_FEATURE

<222> (1)..(44)

<223> Barstar binding barnase-p3 fusion insert

<400> 57

Leu Gln Ser Ser Gly Cys Gly Ser Ser Gly Ser Ser Ile Asn Cys Leu

1 5 10 15

Pro Cys Gly Ala Thr Ser Arg Gly Thr Ser Pro Leu Ala Ser Gly Leu
20, 25 30

Pro Ser Ser Ala Thr Val Gln Cys Leu Ser Ser Ala

35 40

<210> 58

<211> 41

<212> PRT

<213> artificial

<220>

<221> MISC_FEATURE

<222> (1)..(41)

<223> Barstar binding barnase-p3 fusion insert

<400> 58

Leu Gln Ser Ser Gly Lys Ile Val Gln Ala Gly Ala Asn Ile Gln Asp 1 5 10 15

Gly Cys Ile Met His Gly Tyr Cys Asp Thr Asp Thr Ile Val Gly Glu
20 25 30

Asn Gly His Ile Gly Leu Ser Ser Ala

35 40

<210> 59

<211> 45

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(45)

<223> Barstar binding barnase-p3 fusion insert

<400> 59

Leu Gln Ser Ser Gly Val Cys Val Ile Ser Asp Thr Cys Ile Ala Gly

1

5

10

15

Thr Ala Glu Ala Ala Cys Glu Glu Lys Phe Ser Ser Gln Asn Val

20

25

30

Gly His Thr Ile Thr Glu Thr Pro Cys Leu Ser Ser Ala

35

40

45

<210> 60

<211> 44

<212> PRT

<213> artificial

<220>

<221> MISC_FEATURE

<222> (1)..(44)

<223> Barstar binding barnase-p3 fusion insert

<400> 60

Leu Gln Ser Ser Gly Cys Gly Ser Ser Gly Ser Ser Ile Asn Cys Leu

1

5

10

Pro Cys Gly Ala Thr Ser Arg Gly Thr Ser Pro Leu Ala Ser Gly Leu
20 25 30

1.

Pro Ser Ser Ala Thr Ile Gln Cys Leu Ser Ser Ala

35 40

<210> 61

<211> 53

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(53)

<223> Barstar binding barnase-p3 fusion insert

<400> 61

Leu Gln Ser Ser Gly Gln Asp Ser Gln Arg Glu His Ala Ser His Thr

1 5 10 15

Ala Glu Asp Asp Cys Glu Asp Gln Thr Arg Ile His Gln His Ile Arg
20 25 30

Glu Val Asp Phe Val Asp Thr Pro Gln Glu Val Asp Asp Cys Arg Ala

35 40 45

Ala Leu Ser Ser Ala

50

<210> 62

<211> 33

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(33)

<223> Barstar binding barnase-p3 fusion insert

<400> 62

Leu Gln Ser Ser Gly Cys Val Arg Leu Lys Arg Thr Ser Val Asn His

1 10 15

Gln Pro Asp Ala Trp Pro Glu Pro His Leu Lys Ala Ala Cys Glu Pro
20 25 30

Ala

<210> 63

<211> 9

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(9)

<223> Barstar binding barnase-p3 fusion insert

<400> 63

Leu Gln Ser Ser Gly Val Arg Pro Ala

1 5

<210> 64

<211> 44

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(44)

<223> Barstar binding barnase-p3 fusion insert

<400> 64

Leu Gln Ser Ser Gly Cys Gly Ser Ser Gly Ser Ser Ile Asn Cys Leu

1 5 10 15

Pro Cys Gly Ala Thr Ser Arg Gly Thr Ser Pro Leu Ala Ser Gly Leu

20 25 30

Pro Ser Ser Ala Thr Ile Gln Cys Leu Ser Ser Ala

35 40

<210> 65

<211> 30

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(30)

<223> Barstar binding barnase-p3 fusion insert

<400> 65

Leu Gln Ser Ser Gly Thr Glu Val Asp Arg Gly Asn Gln Gln His Asp

1 5 10 15

Thr Asn Asp Arg Asp Phe Thr His Thr Pro Leu Ser Ser Ala

20 25 30

<210> 66

<211> 36

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(36)

<223> Barstar binding barnase-p3 fusion insert

<400> 66

Leu Gln Ser Ser Gly Val Ala Gln Gly Ser Ser Ala Ser Val Asp Val

1 5

15

Thr Ala Thr Asn Ala Val Leu Ser Ala Asp Ser Leu Ser Leu Gly Gly

20 25

30

Gly Glu Pro Ala

35

<210> 67

<211> 19

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(19)

<223> Barstar binding barnase-p3 fusion insert

<400> 67

1

Leu Gln Ser Ser Gly Gly Ala Val Ala Val Thr Pro Gly Pro Val Leu

5

10

15

Ser Ser Ala

<210> 68

<211> 18

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(18)

<223> Barstar binding barnase-p3 fusion insert

<400> 68

Leu Gln Ser Ser Gly His Cys Arg Gly Lys Pro Val Leu Cys Thr His

1

5

10

15

Thr Ala

<210> 69

<211> 9

<212> PRT

<213> Artificial

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<220>
<221> MISC_FEATURE
<222>
      (1)..(9)
<223> Barstar binding barnase-p3 fusion insert
<400> 69
Leu Gln Ser Ser Gly Val Arg Pro Ala
1
                5
<210> 70
<211> 36
<212> PRT
<213> Artificial
<220>
<221> MISC_FEATURE
      (1)..(36)
<222>
<223> Barstar binding barnase-p3 fusion insert
<400> 70
```

. 15

10

Leu Gln Ser Ser Gly Glu Pro Ala Pro Ala His Glu Ala Lys Pro Thr

5

Glu Ala Pro Val Ala Lys Ala Glu Ala Lys Pro Glu Thr Pro Ala His

20 25 30

Leu Ser Ser Ala

35

<210> 71

<211> 33

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(33)

<223> Barstar binding barnase-p3 fusion insert

<400> 71

Leu Gln Ser Ser Gly Cys Val Arg Leu Lys Arg Thr Ser Val Asn His

1 5 10 15

Gln Pro Asp Ala Trp Pro Glu Pro His Leu Lys Ala Ala Cys Glu Pro

20 25 30

Ala

<210> 72

<211> 36

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(36)

<223> Barstar binding barnase-p3 fusion insert

<400> 72

Leu Gln Ser Ser Gly Val Val Asp Trp Ala Lys Met Arg Glu Ile Ala

1 5 10 15

Asp Ser Ile Gly Ala Tyr Leu Phe Val Asp Met Ala His Val Ala Ala
20 25 30

```
Leu Ser Ser Ala
```

35

<210> 73

<211> 117

<212> DNA

<213> Artificial

<220>

<221> misc_feature

<222> (1)..(117)

<223> Vector pK1 polylinker sequence

<400> 73

aatgctggcg gcggcccagc cggcctttct gaggggtcga ctatagaagg acgaggggcc 60

cacgaaggag gtggggtacc cggttccgag ggtggttccg gttccggtga ttttgat 117

<210> 74

<211> 39

<212> PRT

<213> Artificial

<220>

<221> MISC_FEATURE

<222> (1)..(39)

<223> Polypeptide encoded by pK1 vector polylinker sequence

<400> 74

Asn Ala Gly Gly Pro Ala Gly Leu Ser Glu Gly Ser Thr Ile Glu

1 5 10 15

Gly Arg Gly Ala His Glu Gly Gly Gly Val Pro Gly Ser Glu Gly Gly
20 25 30

Ser Gly Ser Gly Asp Phe Asp

35

<210> 75

<211> 117

<212> DNA

<213> Artificial

<220>

<221> misc_feature

<222> (1)..(117)

<223> vector pK2 polylinker sequence

```
<400> 75
aatgctggcg gcggcccagc cggcctttct gaggggtcga ctatagaagg acgagggccc
acgaagcagc tggggtaccg gttccgaggg tggttccggt tccggtgatt ttgatta
<210> 76
<211> 39
<212> PRT
<213> Artificial
<220>
<221> MISC_FEATURE
<222> (1)..(39)
<223> Polypeptide sequence encoded by vector pK2 polylinker region.
<220>
<221> MISC_FEATURE
<222> (38)..(38)
<223> X represents a TGA stop codon
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<221> MISC_FEATURE

<222> (36)..(36)

<223> X represents a stop codon (TGA)

<400> 76

Asn Ala Gly Gly Pro Ala Gly Leu Ser Glu Gly Ser Thr Ile Glu

1 5 10 15

Gly Arg Gly Pro Thr Lys Gln Leu Gly Tyr Arg Phe Arg Gly Trp Phe

20 25 30

Arg Phe Arg Xaa Phe Xaa Leu

35

<210> 77

<211> 35

<212> DNA

<213> Artificial

<220>

<221> misc_feature

<222> (1)..(35)

<223> Sequence of the junction region between Barnase and p3 in recombinant fusion vector fd-3.

<400> 77

atcagactgc aggcggtgcg gccgcagaaa ctgtt

35

<210> 78

<211> 11

<212> PRT

<213> artificial

<220>

<221> MISC_FEATURE

<222> (1)..(11)

<223> Amino acid sequence about the junction of barnase and p3 coding r egions of recombinant fusion vector fd-3.

<400> 78

Ile Arg Leu Gln Ala Ala Ala Glu Thr Val

1

5

10

1

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